3790	2
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### Scientific and Technical Information Center

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SI	EARCH REQUE	ST FORM	
Date: 3/15/S Requester's Full N Art Unit: 1427 Phone (30 Results Format Preferred (circle): PAP	8) 7005	Examiner # Serial Number: 09/2	1. 7779/ 291, 426
To ensure an efficient and quality search, pleas	e attach a copy of the cover s	*************************** heet, claims, and abstract or fill o	**************************************
Title of Invention: Construction	9/ 60-Polyn	n Cibraries	
Inventors (please provide full names):	See A Anch		
Earliest Priority Date: 4/13/			
Search Topic: Please provide a detailed statement of the search elected species or structures, keywords, synonym Define any terms that may have a special meanis	s, acronyms, and registry nun	nbers, and combine with the concep	
*For Sequence Searches Only* Please include a the appropriate serial number.	ll pertinent information (pare	nt, grandchild, divisional, or issue	d patent numbers) along with
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STAFF USE ONLY T	ype of Search  NA Sequence (#)	Vendors and Cost STN	Dialog
Searcher Phone #: 4466	AA Sequence (#)	Questel/Orbit	Dr.Link
Searcher Location:	Structure (#)	Lexis/Nexis	Westlaw
Date Searcher Picked Up: 37 19	Bibliographic	WWW/Internet	
Date Completed: 3   19	Litigation	In-house sequen	ce systems (list)
Searcher Prep & Review Time: 30	Fulltext	Other (specif	y)

# Biotechnology/Chemical Division

Scientific and Technical Information Center



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=> d sta que 180 L38 STR

REP G1=(0-1) AK VAR G2=3/30/38 NODE ATTRIBUTES: DEFAULT MLEVEL IS ATOM DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RSPEC 3 13 24 31
NUMBER OF NODES IS 36

STEREO ATTRIBUTES: NONE

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L41	224	SEA FILE=REGISTRY ABB=ON PLU=ON L40 AND PMS/CI
L42	56	SEA FILE=REGISTRY ABB=ON PLU=ON L41 NOT TYROSIN?
L43	10	SEA FILE=REGISTRY ABB=ON PLU=ON L42 AND ("(C22H23NO6)N" OR
		"(C27H33N06)N" OR "(C26H23N06)N" OR "(C21H21N06)N" OR "(C23H25N
		O6) N" OR "(C25H30N2O5) N" OR "(C25H29NO6) N" OR "(C27H33NO6) N"
		OR "(C23H25NO6)N" OR "(C25H29NO6)N")/MF
L44	168	SEA FILE=REGISTRY ABB=ON PLU=ON L41 NOT L42
L45	178	SEA FILE=REGISTRY ABB=ON PLU=ON (L43 OR L44)
L63	27	SEA FILE=REGISTRY ABB=ON PLU=ON L45 AND (P/ELS OR CLH OR
		C8H12N2O2 OR CH3NO2 OR C6H10O3 OR C34H36N4O7 OR C9H10O3 OR
		CH2O OR C30H28N4O7 OR C8H12NO2O2 OR C44H56N4O7 OR C7H13NO2 OR
		C19H17N3O3 OR C3H7NO2)

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L64
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L65
                                                  L45 AND C24H31NO5 AND
                                          PLU=ON
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L66
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                                                  L45 AND CH2O3 AND C18H19NO5
              1 SEA FILE=REGISTRY ABB=ON
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L67
                AND C20H23NO5
                                                  L45 AND C2H4O
                                          PLU=ON
              6 SEA FILE=REGISTRY ABB=ON
L68
                                          PLU=ON
                                                  L68 AND SQL/FA
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L69
            145 SEA FILE=REGISTRY ABB=ON
                                          PLU=ON
                                                  L45 NOT (L63 OR L64 OR L65
L70
                OR L66 OR L67 OR L69)
                                                   (L63 OR L64 OR L65 OR L66 OR
             38 SEA FILE=REGISTRY ABB=ON
                                          PLU=ON
L71
                L67 OR L68)
              6 SEA FILE=REGISTRY ABB=ON PLU=ON
                                                  L71 AND C2H4O
L72
              5 SEA FILE=REGISTRY ABB=ON
                                          PLU=ON
                                                  L72 NOT SOL/FA
L73
            145 SEA FILE=REGISTRY ABB=ON
                                          PLU=ON
                                                  (L70 OR L73)
L74
            143 SEA FILE=REGISTRY ABB=ON
                                          PLU=ON
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L75
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                                                  L74 NOT L75
              2 SEA FILE=REGISTRY ABB=ON
L76
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                                                  L74 NOT HOMOPOLYMER
            140 SEA FILE=REGISTRY ABB=ON
L77
                                          PLU=ON
                                                  L74 NOT L77
              5 SEA FILE=REGISTRY ABB=ON
L78
                                                  (L76 OR L78)
              6 SEA FILE=REGISTRY ABB=ON
                                           PLU=ON
L79
                                                  L74 NOT L79
            139 SEA FILE=REGISTRY ABB=ON
                                          PLU=ON
L80
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     FILE 'HCAPLUS' ENTERED AT 09:26:13 ON 19 MAR 2001
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E JAMES K/AU
L1
             313 S E3-E15, E27-E44
                 E STEPHEN B/AU
               2 S E4, E5
L2
                 E BROCCHINI S/AU
L3
              36 S E3-E8
               __ E_TANGPASUTHADOL_V/AU____
              11 S E3, E4
L4
                 E KOHN J/AU
             173 S E3, E14-E19
L5
                 E KOEHN J/AU
L6
               3 S E3
                 E KEOHN J/AU
              10 S L1 AND L2-L6
L7
              11 S L2, L3 AND L4-L6
L8
              10 S L4 AND L5, L6
L9
               5 S L7 AND L8, L9
L10
               5 S L8 AND L9
L11
              5 S L10, L11
L12
              16 S L7-L9 NOT L12
L13
              21 S L12, L13
L14
             507 S L1-L7
L15
              52 S L15 AND POLYM?/SC,SX
L16
             128 S L15 AND ?POLYM?
L17
             130 S L16, L17
L18
L19
               7 S L18 AND LIBRARY
               7 S L18 AND COMBINATOR?
L20
              60 S L18 AND ?TYROSIN?
L21
              5 S L21 AND L19, L20
L22
L23
              20 S L14 AND L18
              17 S L23 AND L21
L24
L25
              17 S L22, L24
L26
               4 S L12 AND L25
               1 S L12 NOT L26
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4 S L14 NOT L25

21 S L25-L28

L27

L28

L29

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L30
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              9 S L29 AND POLYESTER
L31
L32
              17 S L30, L31
L33
              21 S L29, L32
                 SEL RN
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L34
L35
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L36
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L37
             130 S L36 AND PMS/CI
L38
                 STR
              37 S L38
L39
L40
          11122 S L38 FUL
                 SAV TEMP L40 HSU291/A
L41
             224 S L40 AND PMS/CI
L42
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L43
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L44
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L45
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L48
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L49
              46 S L47 AND L15
L50
              35 S L49 AND L48
L51
              56 S L48, L50
L52
              42 S L51 NOT P/DT
L53
             11 S L49 NOT L51
L54
              67 S L48-L53
L55
             74 S L33, L54
L56
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L5.7
             18 S L55 NOT L56
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     FILE 'HCAPLUS' ENTERED AT 10:02:49 ON 19 MAR 2001
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L59
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L60
L61
             22 S L55 AND POLYESTER?
L62
             51 S L58-L61
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L63
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L64
              3 S L45 AND (C33H39N3O8 OR C30H32N4O9 OR C32H36N4O10)
L65
              6 S L45 AND C3H7NO2
L66
              1 S L45 AND C24H31NO5 AND C17H19NO3
L67
              1 S L45 AND CH2O3 AND C18H19NO5 AND C20H23NO5
L68
              6 S L45 AND C2H4O
L69
              1 S L68 AND SQL/FA
L70
            145 S L45 NOT L63-L67, L69
L71
             38 S L63-L67, L68
L72
              6 S L71 AND C2H4O
L73
              5 S L72 NOT SQL/FA
L74
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L75
            143 S L74 NOT C32H38N2O7
L76
              2 S L74 NOT L75
            140 S L74 NOT HOMOPOLYMER
L77
L78
              5 S L74 NOT L77
L79
              6 S L76, L78
L80
            139 S L74 NOT L79
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#### SAV L80 HSU291B/A

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L85
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L86
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L87
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L88
              4 S L87 AND L14
L89
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L90
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L91
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L92
              29 S L92 NOT L83
L93
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L94
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                 E JAMES K/AU
L95
              30 S E7-E14
                E BROCHINI S/AU
                E BRODCHINI S/AU
                E BROCCHINI S/AU
              8 S E4-E6
L96
                E STEPHEN S/AU
                E STEPHEN B/AU
                E TANGPASUTHADOL V/AU
                                              L100 = all
                  VARAWUT T/AU
                                              luts for L80, in USPatfull
                E KOHN J/AU
L97
             22 S E10,E11
                E KOEHN J/AU
                E KEOHN J/AU
L98
             13 S L94 AND L95-L97
L99
              3 S L94 NOT L98
L100
             16 S L98, L99
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=> fil uspatful
FILE 'USPATFULL' ENTERED AT 10:45:42 ON 19 MAR 2001
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FILE COVERS 1971 TO PATENT PUBLICATION DATE: 13 Mar 2001 (20010313/PD)
FILE LAST UPDATED: 13 Mar 2001 (20010313/ED)
HIGHEST PATENT NUMBER: US6202212
CA INDEXING IS CURRENT THROUGH 13 Mar 2001 (20010313/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 13 Mar 2001 (20010313/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Dec 2000
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Oct 2000
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>>> is included in file records. A thesaurus is available for the
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>>> fields. This thesaurus includes catchword terms from the
                                                                     <<<
>>> USPTO/MOC subject headings and subheadings. Thesauri are also . <<<
>>> available for the WIPO International Patent Classification
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>>> (IPC) Manuals, editions 1-6, in the /IC1, /IC2, /IC3, /IC4,

>>> /IC5, and /IC (/IC6) fields, respectively. The thesauri in

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>>> the /IC5 and /IC fields include the corresponding catchword <<< >>> terms from the IPC subject headings and subheadings. <<< This file contains CAS Registry Numbers for easy and accurate substance identification. => d 1100 bib abs fhitstr hitrn tot L100 ANSWER 1 OF 16 USPATFULL 2000:124271 USPATFULL Biodegradable, anionic polymers derived from the amino acid L-tyrosine ΤI IN Kohn, Joachim B., Highland Park, NJ, United States Bolikal, Durgadas, Edison, NJ, United States Brode, George L., Bridgewater, NJ, United States Ertel, Sylvie I., Habsheim, France Guan, Shuiyun, Piscataway, NJ, United States Kemnitzer, John E., Plainsboro, NJ, United States PΑ The State University Rutgers, New Brunswick, NJ, United States (U.S. corporation) PΙ US 6120491 20000919 US 1998-56050 19980407 (9) AΙ PRAI US 1997-64656 19971107 (60) Utility DT EXNAM Primary Examiner: Coggins, Wynn Wood; Assistant Examiner: Gring, N. Kent LREP Synnestvedt & Lechner LLP CLMN Number of Claims: 50 ECL Exemplary Claim: 1 DRWN 1 Drawing Figure(s); 1 Drawing Page(s) LN.CNT 1303 CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB A polymer with a hydrolytically labile backbone and having the structure: ##STR1## wherein R.sub.9 is an alkyl, aryl or alkylaryl group with up to 18 carbon atoms having a pendent carboxylic acid group or the benzyl ester thereof; R.sub.12 is an alkyl, aryl or alkylaryl group with up to 18 carbon atoms having a pendent carboxylic acid ester group selected from straight and branched alkyl and alkylaryl esters containing up to 18 carbon atoms and ester derivatives of biologically and pharmaceutically active compounds covalently bonded thereto, provided that the ester group is not a benzyl group or a group that is removed by hydrogenolysis; each R.sub.7 is independently an alkylene group containing up to four carbon atoms; A is selected from: ##STR2## wherein R.sub.8 is selected from saturated and unsaturated, substituted and unsubstituted alkyl, aryl and alkylaryl groups containing up to 18 carbon atoms; k is between about 5 and about 3,000; and  ${\bf x}$  and  ${\bf f}$  independently range from zero to less that one. CAS INDEXING IS AVAILABLE FOR THIS PATENT. 191858-74-9 (1block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles) RN 191858-74-9 USPATFULL CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride and .alpha.-hydro-.omega.-hydroxypoly(oxy-1,2ethanediyl), block (9CI) (CA INDEX NAME)

CRN 133063-33-9 CMF C24 H31 N O5

1

CM

CDES 5:L

Absolute stereochemistry.

CM 2

CRN 25322-68-3

CMF (C2 H4 O)n H2 O

CCI PMS

$$HO - CH_2 - CH_2 - O - I$$

CM 3

CRN 75-44-5 CMF C C12 O

c1-c-c1

IT 191858-74-9

(1block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

IT 191858-68-1P 191858-70-5P

(block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

IT 191858-72-7

(block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

L100 ANSWER 2 OF 16 USPATFULL

AN 2000:105440 USPATFULL

TI Porous polymer scaffolds for tissue engineering

IN Levene, Howard B., Piscataway, NJ, United States

Lhommeau, Christelle M., Highland Park, NJ, United States

Kohn, Joachim B, Highland Park, NJ, United States

PA Rutgers, The State University, New Brunswick, NJ, United States (U.S.

corporation)

PI US 6103255 20000815

AI US 1999-293118 19990416 (9)

DT Utility

EXNAM Primary Examiner: Foelak, Morton

LREP Synnestvedt & Lechner LLP

CLMN Number of Claims: 12

ECL Exemplary Claim: 1

DRWN 1 Drawing Figure(s); 1 Drawing Page(s) LN.CNT 975

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Biodegradable and biocompatible porous scaffolds characterized by a substantially continuous polymer phase, having a highly interconnected bimodal distribution of open pore sizes with rounded large pores of about 50 to about 500 microns in diameter and rounded small pores less than 20 microns in diameter, wherein the small pores are aligned in an orderly linear fashion within the walls of the large pores. Methods of preparing polymeric tissue scaffolds are also disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 188712-99-4P

(porous polymer scaffolds for tissue engineering)

RN 188712-99-4 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, butyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 174702-84-2 CMF C22 H27 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

 ${\tt HO_2C-CH_2-CH_2-CO_2H}$ 

# IT 188712-99-4P 191858-68-1P 219622-84-1P

(porous polymer scaffolds for tissue engineering)

L100 ANSWER 3 OF 16 USPATFULL AN 2000:43762 USPATFULL

TI Copolymers of tyrosine-based polyarlates and poly(alkylene oxides)

IN Kohn, Joachim B., Highland Park, NJ, United States

Yu, Chun, Piscataway, NJ, United States

PA Rutgers, The State University, New Brunswick, NJ, United States (U.S. corporation)

PI US 6048521 20000411

AI US 1998-85571 19980527 (9)

PRAI US 1997-64905 19971107 (60) US 1998-81502 19980413 (60)

US 1997-64656 19971107 (60)

DTUtility

EXNAM Primary Examiner: Kulkosky, Peter F.

LREP Synnestvedt & Lechner LLP

CLMN Number of Claims: 11 ECL Exemplary Claim: 1

6 Drawing Figure(s); 3 Drawing Page(s) DRWN

LN.CNT 917

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Implantable medical devices and drug delivery implants containing AB polyarylate random block copolymers are disclosed, along with methods for drug delivery and for preventing the formation of adhesions between injured tissues employing the polyarylate random block copolymers.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

191858-74-9

(lblock copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

RN 191858-74-9 USPATFULL

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer CN with carbonic dichloride and .alpha.-hydro-.omega.-hydroxypoly(oxy-1,2ethanediyl), block (9CI) (CA INDEX NAME)

CM1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 25322-68-3 CMF (C2 H4 O)n H2 O CCI

$$HO \longrightarrow CH_2 - CH_2 - O \longrightarrow H$$

3 ' CM

CRN 75-44-5 CMF C Cl2 O

#### ΙT 191858-74-9

(1block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

IT 191858-68-1P 191858-70-5P

(block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

ΙT 191858-72-7

> (block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

L100 ANSWER 4 OF 16 USPATFULL 1999:34248 USPATFULL ΑN TI Switch mounting structure IN Yokoyama, Toshiaki, Tokyo, Japan PA Niles Parts Co., Ltd., Japan (non-U.S. corporation) PΤ US 5883348 19990316 US 1997-864361 19970528 (8) ΑI DΤ Utility EXNAM Primary Examiner: Dickson, Paul N.

LREP

Kananen, Ronald P.Rader, Fishman & Grauer

CLMN Number of Claims: 2 ECL Exemplary Claim: 1

3 Drawing Figure(s); 2 Drawing Page(s) DRWN

LN.CNT 326

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A switch mounting structure for fixing integrally, securely and firmly a rotary connector, a switch base, and a bracket mounted to a steering column of a vehicle, by an easy single operation. The switch mounting structure has first frame members 3d having opening portions 3c formed therein provided on a fixing case 3 of a rotary connector 1, and second frame members 7c having opening portions 7b formed therein provided on a bracket 7 of the rotary connector 1. The first frame members 3d are engaged with first elastic engaging portions 5e in first engaging portions 5a provided in a base 5 of a switch 4 by a simple single action. Similarly, the second frame members 7c are engaged with second elastic engaging portions 5h in second engaging portions 5c by a simple single action.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### IT133418-81-2

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

RN 133418-81-2 USPATFULL

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer CN with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

CM 2

CRN 75-44-5 CMF C Cl2 O

### IT 133418-81-2

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

L100 ANSWER 5 OF 16 USPATFULL

AN 1999:27683 USPATFULL

TI Polymeric drug formulations

IN Brocchini, Stephen, Highland Park, NJ, United States Hanson, Stephen R., Stone Mountain, GA, United States Kohn, Joachim B., Highland Park, NJ, United States

PA Rutgers, The State University of New Jersey, New Brunswick, NJ, United States (U.S. corporation)

PI US 5877224 19990302

AI US 1995-508577 19950728 (8)

DT Utility

EXNAM Primary Examiner: Nutter, Nathan M.

LREP Synnestvedt & Lechner CLMN Number of Claims: 13 ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 1312

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Polymeric drug formulations containing a non-releasing single-phase dispersion of a water-soluble drug in a water-insoluble tissue-compatible polymer matrix. Polymeric drug formulations are also disclosed containing a single-phase dispersion of a water-soluble drug and a water-insoluble tissue-compatible polymer matrix, and a second, phase-disrupting polymer that is non-miscible with the tissue-compatible polymer and is present in an amount sufficient to form phase-separated microdomains of the second polymer in the tissue-compatible polymer matrix, so that the release rate of the water-soluble drug from the tissue-compatible polymer matrix is related to the amount of the second polymer. Methods of preparing the polymeric drug formulations are also described, as well as methods for site-specific drug delivery utilizing the polymeric drug formulations.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 149787-39-3

(polymeric drug formulations)

RN 149787-39-3 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with hexanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Me 
$$(CH_2)_5$$
 O O OH

CM 2

CRN 124-04-9 CMF C6 H10 O4

 $HO_2C-(CH_2)_4-CO_2H$ 

### IT 149787-39-3

(polymeric drug formulations)

L100 ANSWER 6 OF 16 USPATFULL

AN 1998:19438 USPATFULL

TI Polymers containing antifibrotic agents, compositions containing such

polymers, and methods of preparation and use
IN Polani, George J. Jamesburg, NJ. United State

Poiani, George J., Jamesburg, NJ, United States Riley, David J., New Brunswick, NJ, United States

Liao, Wei-Chi, Princeton Junction, NJ, United States

Kahn, Joachim, Highland Park, NJ, United States

Gean, Keria Fiorella, Highland Park, NJ, United States

PA University of Medicine & Dentistry of New Jersey, Piscataway, NJ, United States (U.S. corporation)

PI US 5720950 19980224

AI US 1994-260080 19940615 (8)

Pat. No. US 1992-934818, filed on 24 Aug 1992, now patented, Pat. No. US 5372807 which is a continuation-in-part of Ser. No. US 1997-864361, filed on 6 Apr 1997, now abandoned which is a continuation-in-part of Ser. No. US 1991-726301, filed on 5 Jul 1991, now patented, Pat. No. US 5219564 which is a continuation-in-part of Ser. No. US 1990-549494, filed on 6 Jul 1990, now abandoned , said Ser. No. US -864361 which is a continuation of Ser. No. US 1990-523232, filed on 14 May 1990, now abandoned

Utility

EXNAM Primary Examiner: Kulkosky, Peter F.

CLMN Number of Claims: 10

ECL Exemplary Claim: 1

DRWN 17 Drawing Figure(s); 12 Drawing Page(s)

LN.CNT 1861

DT

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention concerns a method for treating fibrotic conditions by administration of an effective amount of an antifibrotic agent. The antifibrotic agent is preferably a proline analog, such as cis-4-hydroxy-L-proline (cHyp). The antifibrotic agent is operatively linked to a monomer or a polymer, with or without a linking compound, e.g., lysine. Intravenous administration is preferred. The present method facilitates the delivery and release of the antifibrotic agent to inhibit collagen accumulation and thereby to treat fibrosis where collagen metabolism is implicated. A reduced quantity of the antifibrotic agent and a corresponding reduction in the potential for toxicity resulting from prolonged administration thereof may be realized.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 133418-81-2

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

RN 133418-81-2 USPATFULL

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer CN with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C C12 O

#### 133418-81-2 ΙT

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

L100 ANSWER 7 OF 16 USPATFULL

ΑN 97:86709 USPATFULL

ΤI Synthesis of tyrosine-derived diphenol monomers

ΙN Kohn, Joachim B., Highland Park, NJ, United States Brocchini, Stephen J., Highland Park, NJ, United States Schwartz, Arthur L., East Windsor, NJ, United States

Rutgers, The State University, New Brunswick, NJ, United States (U.S. PA corporation)

PI

US 5670602 19970923

ΑТ US 1996-625763 19960329 (8)

Division of Ser. No. US 1995-414339, filed on 31 Mar 1995, now patented, RLI Pat. No. US 5587507

DT Utility

EXNAM Primary Examiner: Conrad, Joseph

LREP Lerner, David, Littenberg, Krumholz & Mentlik

CLMN Number of Claims: 7

ECL Exemplary Claim: 1

No Drawings DRWN

LN.CNT 583

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for preparing diphenol compounds, which method includes the AB

steps of coupling a hydroxyphenyl carboxylic acid with a L-tyrosine ester in a water-miscible organic reaction solvent containing a carbodiimide capable of forming a water-soluble urea by-product, thereby forming a diphenol reaction product; and combining the reaction mixture with an amount of water effective to precipitate the diphenol as a water-immiscible organic phase, so that a water-immiscible organic phase is formed containing the diphenol reaction product. New diphenol monomers and polymers polymerized therefrom are also disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 133418-81-2P

(improved prepn. of alkyl desaminotyrosinyltyrosinate as diphenol monomers for polycarbonate polymers)

RN 133418-81-2 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C C12 O

0 || Cl-C-Cl

IT 133418-81-2P 183480-48-0P 183480-50-4P 183480-51-5P 183480-53-7P 183480-54-8P 183480-55-9P

(improved prepn. of alkyl desaminotyrosinyltyrosinate as diphenol monomers for polycarbonate polymers)

L100 ANSWER 8 OF 16 USPATFULL

AN 97:75811 USPATFULL

Polymers containing antifibrotic agents, compositions containing such polymers, and methods of preparation and use

IN Poiani, George J., Jamesburg, NJ, United States
Riley, David J., New Brunswick, NJ, United States
Liao, Wei-Chi, Princeton Junction, NJ, United States
Kahn, Joachim, Highland Park, NJ, United States
Gean, Keria Fiorella, Highland Park, NJ, United States

PA University of Medicine & Dentistry of N.J., Piscataway, NJ, United States (U.S. corporation)

Rutgers University, Piscataway, NJ, United States (U.S. corporation)

PI US 5660822 19970826

AI US 1995-479150 19950607 (8)

Division of Ser. No. US 1994-260080, filed on 15 Jun 1994 which is a continuation-in-part of Ser. No. US 1991-726301, filed on 5 Jul 1991, now patented, Pat. No. US 5219564 76 Ser. No. US 1992-934818, filed on 24 Aug 1992, now patented, Pat. No. US 5372807 which is a continuation-in-part of Ser. No. US 1992-864361, filed on 6 Apr 1992, now abandoned which is a continuation of Ser. No. US 1990-523232, filed on 14 May 1990, now abandoned, said Ser. No. US -726301 which is a continuation of Ser. No. US 1990-549494, filed on 6 Jul 1990, now abandoned

DT Utility

EXNAM Primary Examiner: Kulkosky, Peter F.

LREP Klauber & Jackson CLMN Number of Claims: 13 ECL Exemplary Claim: 1

DRWN 17 Drawing Figure(s); 12 Drawing Page(s)

LN.CNT 1889

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention concerns a method for treating fibrotic conditions by administration of an effective amount of an antifibrotic agent. The antifibrotic agent is preferably a proline analog, such as cis-4-hydroxy-L-proline (cHyp). The antifibrotic agent is operatively linked to a monomer or a polymer, with or without a linking compound, e.g., lysine. Intravenous administration is preferred. The present method facilitates the delivery and release of the antifibrotic agent to inhibit collagen accumulation and thereby to treat fibrosis where collegen metabolism is implicated. A reduced quantity of the antifibrotic agent and a corresponding reduction in the potential for toxicity resulting from prolonged administration thereof may be realized.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 133418-81-2

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

RN 133418=81=2 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C Cl2 O

```
C1-C-C1
```

#### IT 133418-81-2

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

L100 ANSWER 9 OF 16 USPATFULL 97:73701 USPATFULL ΑN Copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) TΤ IN Kohn, Joachim B., Highland Park, NJ, United States Yu, Chun, Piscataway, NJ, United States Rutgers, The State University, New Brunswick, NJ, United States (U.S. PA corporation) PΤ US 5658995 19970819 US 1995-562842 19951127 (8) ΑI DΤ Utility EXNAM Primary Examiner: Mosley, Terressa LREP Lerner, David, Littenberg, Krumholz & Mentlik CLMN Number of Claims: 16 ECL Exemplary Claim: 1 DRWN 6 Drawing Figure(s); 3 Drawing Page(s) CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Random block copolymers having the formula: ##STR1## wherein R.sub.1 is --CH.dbd.CH-- or (--CH.sub.2 --).sub.j, in which j is zero or an integer from one to eight; R.sub.2 is selected from hydrogen, straight and branched alkyl and alkylaryl groups containing up to 18 carbon atoms and derivatives or biologically and pharmaceutically active compounds covalently bonded to said copolymer; each R.sub.3 is independently an alkylene group containing up to 4 carbon atoms; y is an integer between about 5 and about 3000; and f is the percent molar fraction of alkylene oxide in the copolymer and ranges between about 1 and about 99 mole percent. Implantable medical devices and drug delivery implants containing the random block copolymers are also disclosed, along with methods for drug delivery and for preventing the formation of adhesions between injured tissues employing the random block copolymers. Polyarylate random block copolymers are also described.

## CAS INDEXING IS AVAILABLE FOR THIS PATENT.

## IT 191858-74-9

(1block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

RN 191858-74-9 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride and .alpha.-hydro-.omega.-hydroxypoly(oxy-1,2-ethanediyl), block (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

CM 2

CRN 25322-68-3

CMF (C2 H4 O)n H2 O

CCI PMS

$$HO = \begin{bmatrix} CH_2 - CH_2 - O \end{bmatrix}_n$$

CM 3

CRN 75-44-5 CMF C C12 O

IT 191858-74-9

> (1block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

IT191858-68-1P 191858-70-5P

(block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

IT 191858-72-7

(block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

L100 ANSWER 10 OF 16 USPATFULL

96:118718 USPATFULL ΑN

ΤI Synthesis of tyrosine derived diphenol monomers

Kohn, Joachim B., Highland Park, NJ, United States IN Hooper, Kimberly A., Long Valley, NJ, United States

Rutgers, The State University, Piscataway, NJ, United States (U.S. PA

corporation)

ΡI US 5587507 19961224

ΑI US 1995-414339 19950331 (8)

DT Utility

EXNAM Primary Examiner: Conrad, Joseph

LREP Lerner, David, Littenberg, Krumholz & Mentlik

CLMN Number of Claims: 18

ECL Exemplary Claim: 1

No Drawings DRWN

LN.CNT 622

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for preparing diphenol compounds, which method includes the AΒ

steps of coupling a hydroxyphenyl carboxylic acid with a L-tyrosine ester in a water-miscible organic reaction solvent containing a carbodiimide capable of forming a water-soluble urea by-product, thereby forming a diphenol reaction product; and combining the reaction mixture with an amount of water effective to precipitate the diphenol as a water-immiscible organic phase, so that a water-immiscible organic phase is formed containing the diphenol reaction product. New diphenol monomers and polymers polymerized therefrom are also disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 133418-81-2P

(improved prepn. of alkyl desaminotyrosinyltyrosinate as diphenol monomers for polycarbonate polymers)

RN 133418-81-2 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C Cl2 O

PA

IT 133418-81-2P 183480-48-0P 183480-50-4P 183480-51-5P 183480-53-7P 183480-54-8P 183480-55-0P

(improved prepn. of alkyl desaminotyrosinyltyrosinate as diphenol monomers for polycarbonate polymers)

L100 ANSWER 11 OF 16 USPATFULL

AN 95:88245 USPATFULL

TI Poly(alkylene oxide) amino acid copolymers and drug carriers and charged copolymers based thereon

IN Zalipsky, Samuel, Princeton, NJ, United States Bolikal, Durgadas, Edison, NJ, United States Nathan, Aruna, Piscataway, NJ, United States Kohn, Joachim B., Highland Park, NJ, United States

Enzon, Inc., Piscataway, NJ, United States (U.S. corporation)

PI US 5455027 19951003

AI US 1993-23069 19930225 (8)

Pat. No. US 5219564 which is a continuation-in-part of Ser. No. US 1990-549494, filed on 6 Jul 1990, now abandoned

DT Utility

EXNAM Primary Examiner: Webman, Edward J.; Assistant Examiner: Kulkosky, Peter F.

LREP Steinberg, Raskin & Davidson

CLMN Number of Claims: 2 ECL Exemplary Claim: 1

DRWN 2 Drawing Figure(s); 1 Drawing Page(s)

LN.CNT 2251

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Copolymers of poly(alkylene oxides) and amino acids or peptide sequences are disclosed, which amino acids or peptide sequences have pendant functional groups that are capable of being conjugated with pharmaceutically active compounds for drug delivery systems and cross-linked to form polymer matrices functional as hydrogel membranes. The copolymers can also be formed into conductive materials. Methods are also disclosed for preparing the polymers and forming the drug conjugates, hydrogel membranes and conductive materials.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

### IT 133418-81-2

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

RN 133418-81-2 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C Cl2 O

#### IT 133418-81-2

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

L100 ANSWER 12 OF 16 USPATFULL

AN 94:47033 USPATFULL

TI Polyarylates containing derivatives of the natural amino acid 1-tyrosine

Kohn, Joachim B., Highland Park, NJ, United States Fiordeliso, James J., Rochester, NY, United States

Rutgers, The State University of New Jersey, New Brunswick, NJ, United

States (U.S. corporation)

PI US 5317077 19940531

AI US 1993-39929 19930329 (8)

Pat. No. US 5216115 which is a continuation-in-part of Ser. No. US 1991-804767, filed on 9 Dec 1991, now patented, Pat. No. US 5198507 which is a division of Ser. No. US 1990-536425, filed on 12 Jun 1990,

now patented, Pat. No. US 5099060

DT Utility

IN

PA

EXNAM Primary Examiner: Kight, III, John; Assistant Examiner: Mosley, Terressa

LREP Lerner, David, Littenberg, Krumholz & Mentlik

CLMN Number of Claims: 13

ECL Exemplary Claim: 1

DRWN 3 Drawing Figure(s); 2 Drawing Page(s)

LN.CNT 949

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Bioerodible polyarylates derived from biocompatible dicarboxylic acids and natural amino acid-derived diphenol starting materials. Molded articles and controlled drug delivery systems prepared from the polyarylates are also disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 133418-81-2P

(prepn. of biodegradable, for bioprostheses and implants)

RN 133418-81-2 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5

CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C C12 O

IT 133418-81-2P

(prepn. of biodegradable, for bioprostheses and implants)

L100 ANSWER 13 OF 16 USPATFULL

AN 93:48224 USPATFULL

TI Poly(alkylene oxide) amino acid copolymers and drug carriers and charged copolymers based thereon

IN Zalipsky, Samuel, Princeton, NJ, United States Bolikal, Durgadas, Edison, NJ, United States Nathan, Aruna, Piscataway, NJ, United States

Kohn, Joachim B., Highland Park, NJ, United States

PA Enzon, Inc., South Plainfield, NJ, United States (U.S. corporation)

PI US 5219564 19930615

AI US 1991-726301 19910705 (7)

RLI Continuation-in-part of Ser. No. US 1990-549494, filed on 6 Jul 1990,

now abandoned

DT Utility

EXNAM Primary Examiner: Page, Thurman K.; Assistant Examiner: Kulkosky, Peter F.

LREP Lerner, David, Littenberg, Krumholz & Mentlik

CLMN Number of Claims: 20

ECL Exemplary Claim: 1

DRWN 2 Drawing Figure(s); 1 Drawing Page(s)

LN.CNT 2277

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Copolymers of poly(alkylene oxides) and amino acids or peptide sequences are disclosed, which amino acids or peptide sequences have pendant functional groups that are capable of being conjugated with pharmaceutically active compounds for drug delivery systems and cross-linked to form polymer matrices functional as hydrogel membranes. The copolymers can also be formed into conductive materials. Methods are also disclosed for preparing the polymers and forming the drug conjugates, hydrogel membranes and conductive materials.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 133418-81-2

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

RN 133418-81-2 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

Me (CH<sub>2</sub>) 
$$5$$
 O O OH

CM 2

CRN 75-44-5 CMF C C12 O

#### 133418-81-2 TT

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

L100 ANSWER 14 OF 16 USPATFULL

93:44349 USPATFULL ΑN

TΙ Polyarylate containing derivatives of the natural amino acid L-tyrosine

IN Kohn, Joachim B., Highland Park, NJ, United States

Fiordeliso, James J., Rochester, NY, United States

Rutgers, The State University of New Jersey, New Brunswick, NJ, United PA States (U.S. corporation)

US 5216115 19930601 PΤ

ΑI US 1992-930146 19920813 (7)

Continuation-in-part of Ser. No. US 1991-804767, filed on 9 Dec 1991 RLI which is a division of Ser. No. US 1990-536425, filed on 12 Jun 1990, now patented, Pat. No. US 5009060

DT Utility

Primary Examiner: Kight, III, John; Assistant Examiner: Mosley, T. EXNAM

Lerner, David, Littenberg, Krumholz & Mentlik LREP

Number of Claims: 15 CLMN

Exemplary Claim: 1 ECL

3 Drawing Figure(s); 2 Drawing Page(s) DRWN

LN.CNT 945

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Bioerodible polyarylates derived from biocompatible dicarboxylic acids AΒ and natural amino acid-derived diphenol starting materials. Molded articles and controlled drug delivery systems prepared from the polyarylates are also disclosed.

## CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### 149787-38-2P TΤ

(prepn. of, for matrix in controlled drug delivery systems and for medical goods)

RN 149787-38-2 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

CRN 110-15-6 CMF C4 H6 O4

HO2C-CH2-CH2-CO2H

# IT 149787-38-2P 149787-39-3P 149787-40-6P 149787-41-7P 149787-42-8P 149826-02-8P

(prepn. of, for matrix in controlled drug delivery systems and for medical goods)

L100 ANSWER 15 OF 16 USPATFULL

AN 93:24983 USPATFULL

TI Synthesis of amino acid-derived bioerodible polymers

IN Kohn, Joachim B., Highland Park, NJ, United States

Pulapura, Satish K. K., Piscataway, NJ, United States

PA Rutgers, The State University of New Jersey, New Brunswick, NJ, United

States (U.S. corporation)

PI US 5198507 19930330

AI US 1991-804767 19911209 (7)

RLI Division of Ser. No. US 1990-536425, filed on 12 Jun 1990, now patented,

Pat. No. US 5099060

DT Utility

EXNAM Primary Examiner: Buttner, David J.

LREP Lerner, David, Littenberg, Krumholz & Mentlik

CLMN Number of Claims: 7

ECL Exemplary Claim: 1,5

DRWN 1 Drawing Figure(s); 1 Drawing Page(s)

LN.CNT 451

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Novel amino acid-derived polycarbonates and amino acid-derived diphenol compound starting materials from which the polycarbonates are polymerized. Polymer blends of the amino acid-derived polycarbonates with polyiminocarbonates prepared from identical amino acid-derived diphenol starting materials.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 133418-81-2P

(prepn. of biodegradable, for bioprostheses and implants)

RN 133418-81-2 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9

CMF C24 H31 N O5

CDES 5:L

CM2

CRN 75-44-5 CMF C C12 O

#### TΤ 133418-81-2P

(prepn. of biodegradable, for bioprostheses and implants)

L100 ANSWER 16 OF 16 USPATFULL

92:23343 USPATFULL ΑN

ΤI Synthesis of amino acid-derived bioerodible polymers IN Kohn, Joachim B., Highland Park, NJ, United States

Pulapura, Satish K. K., Piscataway, NJ, United States

Rutgers, The State University of New Jersey, New Brunswick, NJ, United PA

States (U.S. corporation) US 5099060 19920324

ΡI

US 1990-536425 19900612 (7) ΑI

DT Utility

Primary Examiner: Gray, Bruce EXNAM

Lerner, David, Littengerg, Krumholz & Mentlik LREP

CLMN Number of Claims: 2

ECL Exemplary Claim: 1

1 Drawing Figure(s); 1 Drawing Page(s) DRWN

LN.CNT 435

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Novel amino acid-derived polycarbonates and amino acid-derived diphenol AΒ compound starting materials from which the polycarbonates are polymerized. Polymer blends of the amino acid-derived polycarbonates with polyiminocarbonates prepared from identical amino acid-derived diphenol starting materials.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### 133418-81-2P ΙT

(prepn. of biodegradable, for bioprostheses and implants)

RN 133418-81-2 USPATFULL

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5

CDES 5:L

CRN 75-44-5 CMF C C12 O

0 || Cl-C-Cl

## IT 133418-81-2P

(prepn. of biodegradable, for bioprostheses and implants)

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L92 ANSWER 1 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 2000:676553 HCAPLUS

DN 134:105786

TI Hydrolytic degradation of tyrosine-derived polycarbonates, a class of new biomaterials. Part of polymeric devices

AU Tangpasuthadol, V.; Pendharkar, S. M.; Peterson, R. C.; Kohn, J.

CS Department of Chemistry, Rutgers-The State University of New Jersey, Piscataway, NJ, 08854, USA

SO Biomaterials (2000), 21(23), 2379-2387 CODEN: BIMADU; ISSN: 0142-9612

PB Elsevier Science Ltd.

DT Journal

LA English

The kinetics and mechanisms of in vitro degrdn. of tyrosine
-derived polycarbonates, a new class of polymeric
biomaterials, were studied extensively at 37.degree.C. These
polymers carry an alkyl ester pendent chain that allows the
fine-tuning of the polymer's material properties, its biol.
interactions with cells and tissue, and its degrdn. behavior. The
polymer carrying an Et ester pendent chain, poly(DTE carbonate),

This set includes
up for L80 + date;
afflicant, all
rep + broader
structure search

has been established as a promising orthopedic implant material, exhibiting bone apposition when in contact with hard tissue. Tyrosine-derived polycarbonates are relatively stable and degrade only very slowly in vitro. Therefore, accelerated studies were conducted at 50 and 65.degree.C to observe the behavior of polymers during the later stages of degrdn. Varying the pendent chain length affected the rate of water uptake, initial degrdn. rate, and phys. stability of the polymeric devices. During the 3-yr study, the polymer degraded by random chain cleavage of the carbonate bonds, accompanied by a relatively small amt. of pendent chain de-esterification. No mass loss was obsd. during this period at 37.degree.C, but mass loss was readily evident during the accelerated studies at 50 and 65.degree.C. Thus, it is reasonable to assume that mass loss will occur also at 37.degree.C, albeit only after extensive backbone carbonate cleavage and pendent chain ester hydrolysis. The dimension and surface area of the devices influenced the initial degrdn. rate, but did not significantly affect the overall rate of degrdn. No evidence of "acid dumping" or the release of acidic residues found during the degrdn. of poly(d, l-lactic acid) were obsd. for this family of tyrosine -derived polycarbonates.

IT 183480-53-7P

RN

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(hydrolytic degrdn. of tyrosine-derived polycarbonates as a class of new biomaterials) 183480-53-7 HCAPLUS

CN Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-(butoxycarbonyl)-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

IT 183480-53-7P 183480-54-8P 319916-60-4P

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(hydrolytic degrdn. of tyrosine-derived polycarbonates as a class of new biomaterials)

RE.CNT 16

(2) Choueka, J; J Biomed Mater Res 1996, V31, P35 HCAPLUS

(3) Ertel, S; J Biomed Mater Res 1994, V28, P919 HCAPLUS

(4) Ertel, S; J Biomed Mater Res 1995, V29(11), P1337 HCAPLUS

(5) Ghorbel, I; J Appl Polym Sci 1995, V55, P173 HCAPLUS

(6) Grizzi, I; Biomaterials 1995, V16(4), P305 HCAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L92 ANSWER 2 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 2000:676552 HCAPLUS

DN 134:105785

TI Hydrolytic degradation of tyrosine-derived polycarbonates, a class of new biomaterials. Part I: Study of model compounds

AU Tangpasuthadol, V.; Pendharkar, S. M.; Kohn, J.

CS Department of Chemistry, Rutgers-The State University of New Jersey, Piscataway, NJ, 08854-8087, USA

SO Biomaterials (2000), 21(23), 2371-2378 CODEN: BIMADU; ISSN: 0142-9612

PB Elsevier Science Ltd.

DT Journal LA English

Tyrosine-derived polycarbonates have been identified AΒ as promising, degradable polymers for use in orthopedic applications. These polymers are non-toxic, biocompatible, and exhibit good bone apposition when in contact with hard tissue. Tyrosine-derived polycarbonates were designed to incorporate two hydrolytically labile bonds in each repeat unit, a carbonate bond that connects the monomer units and an ester bond connecting a pendent chain. The relative hydrolysis rate of the two bonds will det. the type of degrdn. products and the degrdn. pathway of the polymers. In order to study the degrdn. mechanism of these polycarbonates in more detail, a series of small model compds. were designed that mimic the repeat unit of the polymer. Results obtained from the use of these model compds. suggested that the backbone carbonate bond is hydrolyzed at a faster rate than the pendent chain ester bond. Increasing the length of the alkyl pendent chain lowered the hydrolysis rates of both hydrolyzable linkages, possibly by hindering the access of water mols. to those sites. The hydrolysis rates of both linkages were pH dependent with the lowest rate at pH about 5. The results from this study can be used to explain the degrdn. behavior of the corresponding polycarbonates as well as their degrdn. mechanisms. This information is essential when evaluating the utility of tyrosine-derived polycarbonates as degradable medical

implant materials. IT 183480-55-9P

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(hydrolytic degrdn. of tyrosine-derived polycarbonate as a class of biomaterials)

RN 183480-55-9 HCAPLUS

Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-[(octyloxy)carbonyl]-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

## IT 183480-55-9P

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(hydrolytic degrdn. of tyrosine-derived polycarbonate as a class of biomaterials)

RE.CNT 20

RE

CN

- (1) Chasin, M; Biodegradable polymers as drug delivery systems 1990, P43 HCAPLUS
- (2) Choueka, J; J Biomed Mater Res 1996, V31, P35 HCAPLUS
- (4) Doddi, N; US 4052988 1977 HCAPLUS
- (5) Ertel, S; J Biomed Mater Res 1994, V28, P919 HCAPLUS
- (6) Grizzi, I; Biomaterials 1995, V16(4), P305 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
L92
     ANSWER 3 OF 54 HCAPLUS COPYRIGHT 2001 ACS
ΑN
     2000:568454 HCAPLUS
DN
     133:168440
TΙ
     Porous polymer scaffolds for tissue engineering
IN
     Levene, Howard B.; Lhommeau, Christelle M.; Kohn, Joachim B.
PΑ
     Rutgers, the State University, USA
     U.S., 11 pp.
SO
     CODEN: USXXAM
DT
     Patent
     English
LA
FAN.CNT 1
     PATENT NO.
                       KIND DATE
                                             APPLICATION NO.
                                                               DATE
                       ____
                             -----
                                             -----
PΙ
     US 6103255
                        Α
                             20000815
                                             US 1999-293118
                                                               19990416
     WO 2000062829
                       A1
                             20001026
                                             WO 1999-US8375
                                                              19990416
         W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,
             DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS,
              JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,
             MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
              TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
             MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
              ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     AU 9935663
                       A1
                             20001102
                                             AU 1999-35663
                                                               19990416
PRAI US 1999-293118
                       19990416
     WO 1999-US8375
                       19990416
     Biodegradable and biocompatible porous scaffolds characterized by a
AΒ
     substantially continuous polymer phase, having a highly interconnected
     bimodal distribution of open pore sizes with rounded large pores of about
     50 to about 500 .mu.m in diam. and rounded small pores less than 20 .mu.m \,
     in diam., wherein the small pores are aligned in an orderly linear fashion
     within the walls of the large pores. Methods of prepg. polymeric tissue scaffolds are also disclosed. Thus, 0.3 g of poly(L-lactide) was dissolved in 1,4-dioxane/water (91/9% vol./vol.)., the clear soln. was
     then poured on 7 g of sieved sodium chloride salts in a dish. After the
     diffusion of the polymer soln. through the salt bed, it was freeze-dried
     leaving a porous structure. The polymer did not relax during solvent
     removal. Finally, the salt was leached out in water. The water was
     changed several times until the sensitive silver nitrate test did not show
     any addnl. release of chloride ions into the water. The resulting
     scaffolds were removed from the water and dried for several days to const.
          The dried scaffolds were very soft and could be easily deformed
     because of the high total porosity and the low polymer modulus.
ΙT
     188712-99-4P
     RL: DEV (Device component use); PNU (Preparation, unclassified); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
        (porous polymer scaffolds for tissue engineering)
RN
     188712-99-4 HCAPLUS
CN
     L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, butyl ester, polymer
     with butanedioic acid (9CI) (CA INDEX NAME)
     CM
          1
     CRN 174702-84-2
     CMF C22 H27 N O5
     CDES 5:L
```

CM 2

CRN 110-15-6 CMF C4 H6 O4

HO2C-CH2-CH2-CO2H

# IT 188712-99-4P 191858-68-1P 214259-59-3P 219622-84-1P

RL: DEV (Device component use); PNU (Preparation, unclassified); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(porous polymer scaffolds for tissue engineering)

RE.CNT 23

RF.

- (1) Anon; WO 9425079 1994 HCAPLUS
- (3) Cohn; US 4826945 1989 HCAPLUS
- (4) Degroot; Colloid Polym Sci 1990, V268, P1073 HCAPLUS
- (5) Healy; US 5723508 1998 HCAPLUS
- (6) Kohn; US 5099060 1992 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L92 ANSWER 4 OF 54 HCAPLUS COPYRIGHT 2001 ACS
- AN 2000:226537 HCAPLUS
- DN 133:109861
- TI Characterization of the inflammatory response to biomaterials using a rodent air pouch model
- AU Hooper, Kimberly A.; Nickolas, Thomas L.; Yurkow, Edward J.; Kohn, Joachim; Laskin, Debra L.
- CS Department of Pharmacology and Toxicology, Rutgers, The State University of New Jersey, Piscataway, NJ, 08854-8020, USA
- SO J. Biomed. Mater. Res. (2000), 50(3), 365-374 CODEN: JBMRBG; ISSN: 0021-9304
- PB John Wiley & Sons, Inc.
- DT Journal
- LA English
- Using a rodent air pouch, the inflammatory responses to biomaterials with distinct phys. properties and chem. compns. were compared. The polymers examd. were expanded poly(tetrafluoroethylene) (ePTFE), silicone, low-d. polyethylene (LDPE), poly(L-lactic acid) (PLLA), poly(desaminotyrosyltyrosine Et carbonate) [poly(DTE carbonate)], and poly(desaminotyrosyltyrosine benzyl carbonate) [poly(DTBzl carbonate)]. We found that implantation of disks (4.5-4.8 mm) of these materials into rodent air pouches for 2 days had no effect on the no. or type of cells recovered relative to sham controls. With each of the materials, macrophages were the predominant cell type identified (60-75%), followed by granulocytes (20-25%) and lymphocytes (10%). Implantation of poly(DTE carbonate),

ePTFE, LDPE, or poly(DTBzl carbonate) into the pouches for 2 days caused an increase in release of superoxide anion by the pouch cells. Cells from pouches contg. poly(DTE carbonate) also released more hydrogen peroxide and were more phagocytic. In contrast, PLLA and silicone had no effect on the functional activity of cells recovered from the pouches. Prolonging the implantation time of poly(DTE carbonate) or PLLA to 7 days did not alter the no. or type of cells isolated from the pouches. However, cells from pouches contg. poly(DTE carbonate) for 7 days continued to produce increased quantities of superoxide anion relative to sham control pouch cells. These results suggest that the air pouch model is a highly sensitive method and therefore useful for evaluating the functional responses of inflammatory cells to biomaterials.

IT 219622-84-1

RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(characterization of inflammatory response to biomaterials using rodent air pouch model)

RN 219622-84-1 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, ethyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 135313-59-6 CMF C20 H23 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C Cl2 O

## IT 219622-84-1 219622-85-2

RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(characterization of inflammatory response to biomaterials using rodent air pouch model)

RE.CNT 41

RE

- (2) Anderson, J; Biomaterials 1984, V5, P5 HCAPLUS
- (3) Athanasiou, K; Biomaterials 1996, V17, P93 HCAPLUS
- (6) Behling, C; J Biomed Mater Res 1986, V20, P653 HCAPLUS
- (7) Bergsma, J; J Biomed Mater Res 1995, V29, P173 HCAPLUS
- (9) Cerami, A; Clin Immunol Immunopath 1992, V62, PS3 HCAPLUS ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L92 ANSWER 5 OF 54 HCAPLUS COPYRIGHT 2001 ACS

```
2000:225757 HCAPLUS
ΑN
DN
     132:348250
TТ
     Characterization of combinatorially designed polyarylates by
     time-of-flight secondary ion mass spectrometry
ΑU
     Belu, Anna M.; Brocchini, Stephen; Kohn, Joachim;
     Ratner, Buddy D.
CS
     Department of Bioengineering, University of Washington, Seattle, WA,
     98195-1750, USA
SO
     Rapid Commun. Mass Spectrom. (2000), 14(7), 564-571
     CODEN: RCMSEF; ISSN: 0951-4198
PB
     John Wiley & Sons Ltd.
DT
     Journal
LA
     English
     A series of 16 polyarylates, with well-controlled and systematically
AB
     varying chem., has been characterized by time-of-flight secondary ion mass
     spectrometry (TOF-SIMS). The polymers are structurally
     identical except for the incremental addns. of C2H4 units to the backbone
     and side-chain. From the spectra, peaks characteristic of all
     polyarylates are identified. Furthermore, evaluation of the spectra and
     identification of unique signals allow classification of the polyarylates
     according to side-chain and backbone chem.
IT
     149787-38-2
     RL: PRP (Properties)
        (time-of-flight secondary ion mass spectrometry of tyrosine
        ester-based polyesters)
RN
     149787-38-2 HCAPLUS
CN
     L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer
     with butanedioic acid (9CI) (CA INDEX NAME)
    CM
          1
    CRN
         133063-33-9
    CMF C24 H31 N O5
    CDES 5:L
```

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

 ${\tt HO_2C-CH_2-CH_2-CO_2H}$ 

IT 149787-38-2 149787-39-3 149787-40-6 149787-41-7 149826-02-8 188712-92-7 188712-95-0 188712-97-2 188712-99-4 188713-01-1 188713-03-3 188713-05-5 188713-08-8 188713-09-9 188713-10-2 188713-11-3 RL: PRP (Properties) (time-of-flight secondary ion mass spectrometry of tyrosine
ester-based polyesters)

RE.CNT 5

RE

(1) Brocchini, S; J Am Chem Soc 1997, V119, P4553 HCAPLUS

- (2) Fiordeliso, J; J Biomater Sci Polym Ed 1994, V5, P497 HCAPLUS
- (3) Perez-Luna, V; J App Polym Sci 1997, V63, P1467 HCAPLUS
- (4) Reichlmaier, S; Surf Interface Anal 1994, V21, P739 HCAPLUS
- (5) Tamada, Y; J Biomed Mater Res 1994, V28, P283
- L92 ANSWER 6 OF 54 HCAPLUS COPYRIGHT 2001 ACS
- AN 2000:108769 HCAPLUS
- DN 132:270023
- TI PEG-variant biomaterials as selectively adhesive protein templates: model surfaces for controlled cell adhesion and migration
- AU Tziampazis, Evangelos; Kohn, Joachim; Moghe, Prabhas V.
- CS Department of Chemical and Biochemical Engineering, Rutgers University, Piscataway, NJ, 08854-8058, USA
- SO Biomaterials (2000), 21(5), 511-520 CODEN: BIMADU; ISSN: 0142-9612
- PB Elsevier Science Ltd.
- DT Journal
- LA English
- AΒ Our study focused on the role of poly(ethylene glycol) (PEG) in actively regulating the biol. responsiveness of protein-adsorbed biomaterials. To this end, we designed PEG-variant biomaterials from a family of tyrosine/PEG-derived polycarbonates to present surfaces ranging from low to intermediate levels of PEG concn., below the PEG level requisite for complete abolition of protein adsorption. We analyzed the effect of PEG concn. on the amt., conformation and bioactivity of an adsorbed model protein, fibronectin, and on the attachment, adhesion strength and motility of L929 fibroblasts. Our results demonstrate that low levels of PEG can regulate not only the extent but also the conformation and specific bioactivity of adsorbed fibronectin. As the PEG concn. was increased from 0 to 6 mol%, the amt. of adsorbed fibronectin decreased linearly yet the fibronectin conformation was altered such that the overall bioactivity of adsorbed fibronectin was uncompromised. We report that the degree of cell attachment varied with PEG concn. in a manner similar to the dependence of fibronectin bioactivity on PEG. In contrast, the nature of cell adhesion strength dependence on PEG paralleled the pattern obsd. for fibronectin surface concn. Our studies also indicated that the rate of cell migration was inversely correlated with PEG concn. over a narrow range of PEG concn. Overall, these results highlight the striking ability of PEG-variant biomaterials to systematically regulate the behavior of adsorbed cell adhesion proteins and, consequently, effect cell functions.

IT 263565-88-4

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (PEG-variant biomaterials as selectively adhesive protein templates as model surfaces for controlled cell adhesion and migration)

RN 263565-88-4 HCAPLUS

CM 1

CRN 135313-59-6 CMF C20 H23 N O5 CDES 5:L

CM 2

CRN 85022-96-4

CMF (C2 H4 O)n C2 H2 O5

CCI PMS

$$HO_2C$$
  $O-CH_2-CH_2$   $O-CO_2H$ 

#### IT 263565-88-4

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(PEG-variant biomaterials as selectively adhesive protein templates as model surfaces for centralled cell adhesive and migration)

model surfaces for controlled cell adhesion and migration)

RE.CNT 49

RE

- (1) Absolom, D; Biochim Biophys Acta 1981, V670, P74 HCAPLUS
- (2) Absolom, D; J Biomed Mater Res 1987, V21, P161 HCAPLUS
- (3) Amiji, M; Biomaterials 1992, V13, P682 HCAPLUS
- (4) Andrade, J; Interfacial phenomena and bioproducts 1996, P19 HCAPLUS
- (5) Arakawa, T; Biochemistry 1985, V24, P6756 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L92 ANSWER 7 OF 54 HCAPLUS COPYRIGHT 2001 ACS
- AN 2000:5405 HCAPLUS
- DN 132:212631
- TI Small changes in **polymer** chemistry have a large effect on the bone-implant interface: evaluation of a series of degradable **tyrosine**-derived **polycarbonates** in bone defects
- AU James, Kenneth; Levene, Howard; Parsons, J. Russell; Kohn, Joachim
- CS Department of Chemistry, Rutgers, The State University of New Jersey, New Brunswick, NJ, 09803, USA
- SO Biomaterials (1999), 20(23/24), 2203-2212 CODEN: BIMADU; ISSN: 0142-9612
- PB Elsevier Science Ltd.
- DT Journal
- LA English

AB

In a series of homologous, tyrosine-based polycarbonates, small changes in the chem. structure of the polymer pendent chain affectes the bone response in a long-term (1280 days) implantation study. Identically sized pins, prepd. from poly(DTE carbonate), poly(DTB carbonate), poly(DTH carbonate), and poly(DTO carbonate) were implanted transcortically in the proximal tibia and the distal femur of skeletally mature New Zealand White Rabbits. The tissue response at the bone-implant interface was characterized in terms of the absence of a fibrous capsule (direct bone apposition, indicative of a bone bonding response) or the presence of a fibrous capsule (referred to as the encapsulation response). The relative frequency of direct bone apposition vs. encapsulation was recorded for each polymer throughout the entire period of the study. While all 4 polymers were tissue compatible, there was a

correlation between the chem. structure of the pendent chain and the type of bone response obsd., with poly(DTE carbonate) having the highest tendency to elicit direct bone apposition. Based on in vivo degrdn. data and the ability of model polymers with carboxylate groups at their surface to chelate calcium ions, it is proposed that the ability of poly(DTE carbonate) to bond to bone is caused by the facile hydrolysis of the pendent Et ester groups which creates calcium ion chelation sites on the polymer surface. The incorporation of calcium chelation sites into the chem. structure of an implant material appears to be a key requirement if direct bone apposition/bone bonding is desired. Very subtle changes in the chem. compn. of an implant material can have significant effects on the long-term tissue response in a clin. relevant model.

ΙT 183480-53-7

> RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(degradable tyrosine-derived polycarbonates evaluation in bone defects)

RN183480-53-7 HCAPLUS

> Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-(butoxycarbonyl)-1,2ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

IT 183480-53-7 183480-54-8 183480-55-9

> RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(degradable tyrosine-derived polycarbonates evaluation in bone defects)

RE.CNT

RE

CN

- (2) Boyan, B; Biomaterials 1996, V17, P137 HCAPLUS
- (3) Brocchini, S; J Amer Chem Soc 1997, V119(19), P4553 HCAPLUS
- (4) Brocchini, S; J Biomed Mater Res 1998, V42, P66 HCAPLUS
- (5) Choueka, J; J Biomed Mater Res 1996, V31, P35 HCAPLUS
- (7) Ertel, S; J Biomed Mater Res 1994, V28, P919 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L92 ANSWER 8 OF 54 HCAPLUS COPYRIGHT 2001 ACS

ΑN 1999:784854 HCAPLUS

DN 132:93927

TI Conductivity and high-temperature relaxation of tyrosine-derived polyarylates measured with thermal stimulated currents

ΑU Puma, M.; Suarez, N.; Kohn, J.

- CS ENINSEL, Carr. Nac. Hoyo de la Puerta, Instituto de Ingenieria, Sartenejas-Caracas, Venez.
- J. Polym. Sci., Part B: Polym. Phys. (1999), 37(24), 3504-3511 SO CODEN: JPBPEM; ISSN: 0887-6266
- PB John Wiley & Sons, Inc.
- DTJournal
- LA English
- AB Thermal-stimulated polarization and depolarization expts. without blocking electrodes are performed on tyrosine-derived polyarylates with different backbone lengths. The expts. on the different samples are carried out using the same thermal history throughout the entire characterization process. The high-temp. current rise caused by the cond. of the samples

is studied with a simple model that utilizes an approxn. of the Williams-Landel-Ferry (WLF) relaxation time. The cond. data is well reproduced except for temps. well below the glass-transition temp. and for small currents. The glass-transition peak is modeled with a phenomenol. expression valid near Tg, which is able to describe the glass relaxation with a min. no. of parameters. The conduction and the glass-transition relaxation are studied vs. the structural changes for the different samples. It is found that the cond. and the glass-transition temp. shift to lower temps. as the methylene groups in the backbone increase. Furthermore, if the exptl. data is presented as a function of the reduced temp., the shape of the glass-transition relaxation for the different samples is independent of the polymer backbone length.

IT 149787-38-2

RL: PRP (Properties)

(cond. and high-temp. relaxation of tyrosine-derived polyarylates measured with thermal stimulated currents)

RN 149787-38-2 HCAPLUS CN

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

CM

133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

2 CM

CRN 110-15-6 CMF C4 H6 O4

 ${\rm HO_2C-CH_2-CH_2-CO_2H}$ 

IT 149787-38-2 149787-39-3 149787-40-6 188713-08-8

RL: PRP (Properties)

(cond. and high-temp. relaxation of tyrosine-derived polyarylates measured with thermal stimulated currents)

RE.CNT

RE

- (3) Brocchini, S; J Biomed Mater Res 1998, V42, P66 HCAPLUS
- (4) Canadas, J; J Polymer 1998, V39, P2795 HCAPLUS
- (5) Chee, K; J Appl Polym Sci 1987, V33, P1067 HCAPLUS
- (7) Fiordeliso, J; J Biomater Sci Polym Ed 1994, V5, P497 HCAPLUS
- (9) Kohn, J; US 5216115 1993 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT

L92 ANSWER 9 OF 54 HCAPLUS COPYRIGHT 2001 ACS

ΑN 1999:722402 HCAPLUS

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DN 132:313446
```

TI Polymer ordering enhances delivery of antithrombotic peptide

AU Schachter, D.; Kohn, J.

- CS Department of Chemistry, Rutgers University, Piscataway, NJ, 08855-0939, USA
- SO Proc. Int. Symp. Controlled Release Bioact. Mater. (1999), 26th, 625-626 CODEN: PCRMEY; ISSN: 1022-0178
- PB Controlled Release Society, Inc.

DT Journal

LA English

AB Strong peptide-polymer interactions can occur when formulating a peptide into tyrosine-derived polyarylates because of the peptide-like structure. In the case of polymers that end to order, these interactions remaine strong only in the amorphous regions. In the cryst. regions, the self assocn. of the polymer chains excludes these interactions allowing diffusion to occur.

IT 149787-39-3

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(polymer ordering enhances delivery of antithrombotic peptide)

RN 149787-39-3 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with hexanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

Me 
$$(CH_2)$$
  $5$   $O$   $O$   $OH$   $HO$ 

CM 2

CRN 124-04-9 CMF C6 H10 O4

 $HO_2C^-$  (CH<sub>2</sub>)<sub>4</sub> - CO<sub>2</sub>H

## IT 149787-39-3 149826-02-8 188712-97-2 188713-11-3

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(polymer ordering enhances delivery of antithrombotic peptide)

RE.CNT 5

RE

- (1) Brocchini, S; JACS 1997, V119, P19
- (2) Gombotz, W; Bioconjugate Chem 1995, V6 HCAPLUS
- (3) Schachter, D; Proc of CRS Conf on Adv in Control 1996
- (4) Schachter, D; manuscript in preparation
- (5) Tcheng, J; Circulation 1995, V91, P8

```
ANSWER 10 OF 54 HCAPLUS COPYRIGHT 2001 ACS
L92
     1999:672896 HCAPLUS
AN
DN
     131:299854
ΤI
     Copolymer libraries, their production and monomers
     therefor
     Kohn, Joachim B.; Brocchini, Stephen; James,
IN
     Kenneth; Tangpasuthadol, Varawut
     Rutgers, the State University, USA
PΑ
SO
     PCT Int. Appl., 69 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 3
     PATENT NO.
                        KIND
                               DATE
                                               APPLICATION NO.
     WO 9952962
                         A1
                               19991021
                                               WO 1999-US8131
                                                                  19990413 <--
PΙ
          W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,
              DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
              MD, RU,
                       TJ, TM
          RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
              ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
                      GA, GN, GW, ML, MR, NE, SN, TD, TG
              CI, CM,
                                              AU 1999-36420
                                                                  19990413 <--
     AU 9936420
                               19991101
                         Α1
                                               EP 1999-918534
                                                                19990413 <--
                               20010207
     EP 1073688
                         Α1
              AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, FI
PRAI US 1998-81502
                        19980413 <--
     WO 1999-US8131
                        19990413
     A multidimensional copolymer array of a plurality of
AΒ
     copolymers, polymd. from at least two independently
     variable sets of monomers, is created. The homologous variations of the
     monomer series are selected to det. the effect of varying the structural
     features of the copolymer on at least one end-use property of
     the copolymer. Methods for detg. the effects as a function of
     variation within the monomer series and identifying members having useful
     properties are also disclosed. In an example, a library of 112
     polyesters is obtained from parallel syntheses involving 9
     different dicarboxylic acids and 14 different amide linkage-contg.
     bisphenols (produced by condensing L-tyrosine esters with
     4-hydroxyphenylacetic or -propionic acid). The polyamide-
     polyesters show good biocompatibility and the synthesis method
     facilitates the selection of polymers for biomedical
     applications.
ΙT
     149787-38-2P
     RL: BAC (Biological activity or effector, except adverse); IMF (Industrial
     manufacture); THU (Therapeutic use); BIOL (Biological study); PREP
      (Preparation); USES (Uses)
         (prodn. of biocompatible polyamide-polyester
      libraries)
RN
     149787-38-2 HCAPLUS
     L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer
CN
     with butanedioic acid (9CI) (CA INDEX NAME)
     CM
          133063-33-9
     CRN
     CMF
          C24 H31 N O5
     CDES 5:L
```

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

HO2C-CH2-CH2-CO2H

libraries)

```
IT
    149787-38-2P 149787-39-3P 149787-40-6P
    149787-41-7P 149826-02-8P 188712-92-7P
    188712-95-0P 188712-97-2P 188712-99-4P
    188713-01-1P 188713-03-3P 188713-05-5P
    188713-08-8P 188713-09-9P 188713-10-2P
    188713-11-3P 189760-07-4P 189760-09-6P
    189760-11-0P 189760-12-1P 189760-14-3P
    189760-16-5P 247076-59-1P 247076-60-4P
    247076-61-5P 247076-62-6P 247076-63-7P
    247076-64-8P 247076-65-9P 247076-66-0P
    247076-67-1P 247076-68-2P 247076-69-3P
    247076-70-6P 247076-71-7P 247076-72-8P
    247076-73-9P 247076-74-0P 247076-75-1P
     247076-76-2P 247076-77-3P 247076-78-4P
     247076-79-5P 247076-80-8P 247076-81-9P
    247076-82-0P 247076-83-1P 247076-84-2P
    247076-85-3P 247076-86-4P 247076-87-5P
    247076-88-6P 247076-89-7P 247076-90-0P
    247076-92-2P 247076-94-4P 247076-96-6P
    247076-98-8P 247077-00-5P 247077-03-8P
    247077-05-0P 247077-07-2P 247077-09-4P
    247077-10-7P 247077-12-9P 247077-14-1P
    247077-16-3P 247077-18-5P 247077-20-9P
    247077-22-1P 247077-24-3P 247077-26-5P
    247077-28-7P 247077-31-2P 247077-33-4P
    247077-35-6P 247077-37-8P 247077-39-0P
    247077-41-4P 247077-43-6P 247077-45-8P
    247077-47-0P 247077-48-1P 247077-49-2P
    247077-50-5P 247077-52-7P 247077-54-9P
     247077-56-1P 247077-57-2P 247077-58-3P
     247077-59-4P 247077-60-7P 247077-61-8P
     247077-62-9P 247077-63-0P 247077-64-1P
     247077-65-2P 247077-66-3P 247077-67-4P
     247077-68-5P 247077-69-6P 247077-70-9P
     247077-71-0P 247077-72-1P 247077-73-2P
     247077-74-3P 247077-75-4P 247077-76-5P
     247077-77-6P 247077-78-7P 247077-79-8P
     247077-80-1P
     RL: BAC (Biological activity or effector, except adverse); IMF (Industrial
     manufacture); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (prodn. of biocompatible polyamide-polyester
```

RE.CNT 4

RE

- (1) Brandstetter; US 5587423 A 1996 HCAPLUS
- (2) Brocchini; J Am Chem Soc 1997, V119, P4553 HCAPLUS
- (3) Gravert; 214th ACS National Meeting, Parallel Polymer Synthesis: A Novel Bifunctional Radical Initiator that Allows Discrete Combinatorial Polymer Synthesis 1997, Part 2
- (4) Gravert; Bifunctional Initiators for Free Radical Polymerization of Non-crosslinked Block Copolymers. Tet Letters 1998, V39, P1513 HCAPLUS
- L92 ANSWER 11 OF 54 HCAPLUS COPYRIGHT 2001 ACS
- AN 1999:146214 HCAPLUS
- TI The use of combinatorial approaches in the design of new biomaterials
- AU Kohn, J.; Brocchini, S.; James, K.; Tangpasuthadol, V.
- CS Department of Chemistry, Rutgers University, Piscataway, NJ, 08854, USA
- SO Book of Abstracts, 217th ACS National Meeting, Anaheim, Calif., March 21-25 (1999), POLY-178 Publisher: American Chemical Society, Washington, D. C.

CODEN: 67GHA6

- DT Conference; Meeting Abstract
- LA English
- AB The development of combinatorial approaches to polymer design provides new strategies for the rapid identification of novel biomaterials. Our prior experience with the design of polymeric biomaterials indicated that the traditional methods of synthesizing new polymers one after the other in a sequential fashion are too time consuming to satisfy the materials need of emerging fields such as "tissue engineering". To address this problem, we have created the first combinatorial library of biomaterials. In this library of 112 structurally related polymers, it has been possible to identify new correlations between polymer structure, polymer properties, and the cellular response in vitro. Data will be presented that illustrate the value of this combinatorial approach in identifying useful new polymeric compns. for a range of medical applications.
- L92 ANSWER 12 OF 54 HCAPLUS COPYRIGHT 2001 ACS
- AN 1999:109959 HCAPLUS
- DN 130:297273
- TI Tyrosine-PEG-derived poly(ether carbonate)s as new biomaterials. Part II: study of inverse temperature transitions
- AU Yu, Chun; Mielewczyk, Slawomir S.; Breslauer, Kenneth J.; Kohn, Joachim
- CS Department of Chemistry, Rutgers, The State University of New Jersey, New Brunswick, NJ, 08903, USA
- SO Biomaterials (1999), 20(3), 265-272 CODEN: BIMADU; ISSN: 0142-9612
- PB Elsevier Science Ltd.
- DT Journal
- LA English
- Tyrosine-poly(alkylene oxide)-derived poly(ether carbonates) represent a new group of degradable biomaterials that exhibit inverse temp. transitions. Poly(DTE co 70%PEG1000 carbonate) (DTE = deaminotyrosyltyrosine Et ester) was chosen as an example to study this special phase transition behavior of the polymers. The obsd. transition temp. varied slightly depending on the technique used, e.g. CD spectra always gave a lower temp. than UV/visible spectra. The CD and UV/visible studies indicated that the transition temp. was both heating rate and concn. dependent. Thermodn. parameters of the transition (enthalpy, entropy, and free energy) were detd. by DSC. The molecularity of the transition was 2.6, as calcd. from UV and DSC data. The transition temp. could be varied from 18 to 580C by changing the polymer structure.
- IT 223114-10-1
  - RL: PRP (Properties)
  - (inverse temp. phase transition properties of biodegradable)
- RN 223114-10-1 HCAPLUS
- IT 223114-10-1

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RL: PRP (Properties)
        (inverse temp. phase transition properties of biodegradable)
RE.CNT
       18
RE
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(3) Chen, G; Nature 1995, V373, P49 HCAPLUS
(4) Cheng, Y; Macromolecules 1995, V28(8), P2665 HCAPLUS
(5) Fujishige, S; J Phys Chem 1989, V93, P3311 HCAPLUS
(6) Hirotsu, S; J Chem Phys 1987, V87(2), P1392 HCAPLUS
ALL CITATIONS AVAILABLE IN THE RE FORMAT
L92
     ANSWER 13 OF 54 HCAPLUS COPYRIGHT 2001 ACS
AN
     1999:109957 HCAPLUS
DN
     130:297251
TΙ
     Tyrosine-PEG-derived poly(ether carbonate)s as new biomaterials part. I:
     synthesis and evaluation
ΑU
     Yu, Chun; Kohn, Joachim
CS
     Department of Chemistry, Rutgers, The State University of New Jersey, New
     Brunswick, NJ, 08903, USA
SQ
     Biomaterials (1999), 20(3), 253-264
     CODEN: BIMADU; ISSN: 0142-9612
PB
     Elsevier Science Ltd.
DT
     Journal
LA
     English
AB
     Tyrosine-PEG-derived poly(ether carbonates) were prepd. by condensation
     copolymn. of PEG and deaminotyrosyltyrosine alkyl esters with phosgene.
     The resulting polymers were random copolymers with wt.-av. mol. wts.
     40,000-200,000. Chem. structure and purity were confirmed by NMR and FTIR
     spectral anal. General structure-property correlations were established.
     The glass transition temp. decreased with increasing PEG content and
     increasing alkyl ester chain length. When higher mol. wt. PEG blocks were
     used, the glass transition temp. increased relative to identical polymers
     having shorter PEG blocks. The tensile modulus increased with decreasing
     PEG content, decreasing pendent chain length, and when longer PEG blocks
     were used. Water uptake and the rate of backbone degrdn. increased with
     increasing PEG content. Microspheres could be prepd. by solvent evapn.
     techniques from copolymers with low PEG content. Release rate of
     p-nitroaniline and fluorescein isothiocyanate-dextran from the
    microspheres increased with increasing PEG content. While
     tyrosine-derived polycarbonates were excellent substrates for cell
     attachment and growth, the presence of only 5 mol% of PEG1000 led to low
     or no cell attachment in short-term cell culture with both rat lung
     fibroblasts and osteoblasts. The polymers were non-cytotoxic.
IT
     223114-10-1P
     RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (prepn. and properties of tyrosine-based polyoxyethylene-
        polycarbonates)
RN
     223114-10-1 HCAPLUS
     223114-10-1P 223114-11-2P 223114-13-4P
IT
     223114-15-6P
     RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (prepn. and properties of tyrosine-based polyoxyethylene-
       polycarbonates)
RE.CNT
       40
RE
(1) Bakker, D; J Biomed Mater Res 1990, V24(4), P489 HCAPLUS
(5) Brocchini, S; J Amer Chem Soc 1997, V119(19), P4553 HCAPLUS
(6) Cho, C; J Biomed Mater Res 1993, V27, P199 HCAPLUS
(7) Choueka, J; J Biomed Mater Res 1996, V31, P35 HCAPLUS
(9) Engelberg, I; Biomaterials 1991, V12(3), P292 HCAPLUS
ALL CITATIONS AVAILABLE IN THE RE FORMAT
L92
    ANSWER 14 OF 54 HCAPLUS COPYRIGHT 2001 ACS
```

1999:34955 HCAPLUS

ΑN

```
DN
      130:110770
      Biphasic polymerization process using hydrolysis-sensitive monomers
ΤI
ΙN
      Kemnitzer, John E.; Brode, George L.; Kohn, Joachim B.
      Integra Lifesciences I, Ltd., USA
PA
      PCT Int. Appl., 39 pp.
SO
      CODEN: PIXXD2
DT
      Patent
      English
LA
FAN.CNT 1
      PATENT NO.
                         KIND DATE
                                                  APPLICATION NO.
                                                                    DATE
                                -----
                                                  -----
      ______
                         ____
      WO 9900442
                         A1
                                19990107
                                                 WO 1998-US13657 19980629 <--
PΪ
          W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG,
               KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
          NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG
      AU 9881786
                                 19990119
                                                  AU 1998-81786
                                                                      19980629 <--
                          A1
                                 20000412
                                                  EP 1998-931748
      EP 991693
                          A1
                                                                      19980629 <--
          R: AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, NL, SE, IE, FI
PRAI US 1997-884108
                         19970627
     WO 1998-US13657
                         19980629
AB
      Improvements are disclosed for biphasic polymn. processes in which an aq.
      soln. of a first monomer that is hydrolytically unstable below a pH of
      about six or above a pH of about eight is admixed with a water-immiscible
      org. solvent and there is added to the admixt. a catalyst selected from
      tertiary amine, quaternary amine and phosphonium catalysts, an
      acid-forming co-monomer for the first monomer, an acid scavenger, after
     which the resulting polymer is recovered, wherein the improvement includes
     providing the aq. soln. at a pH between about six and about eight, and
     adding to the admixt. the acid-forming co-monomer and the acid scavenger
     at relative rates effective to maintain the pH of the admixt. between about six and about eight. The catalyst may be added in a molar ratio to
     the first monomer effective to provide a predetd. wt.-av. or no.-av. mol.
     wt. for the resulting polymer. Biphasic polymn. processes for monomers
     that are not pH sensitive are also disclosed.
ΙT
     219622-84-1P, Desaminotyrosyltyrosine ethyl ester-phosgene
     copolymer
     RL: IMF (Industrial manufacture); PREP (Preparation)
         (biphasic polymn. process using hydrolysis-sensitive monomers)
RN
     219622-84-1 HCAPLUS
CN
     L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, ethyl ester, polymer
     with carbonic dichloride (9CI) (CA INDEX NAME)
     CM
          135313-59-6
     CRN
     CMF C20 H23 N O5
     CDES 5:L
```

Absolute stereochemistry.

```
CM
          2
     CRN 75-44-5
     CMF C Cl2 O
C1-C-C1
     219622-84-1P, Desaminotyrosyltyrosine ethyl ester-phosgene
IT
     copolymer 219622-85-2P, Desaminotyrosyltyrosine benzyl
     ester-phosgene copolymer 219622-86-3P 219638-85-4P
     RL: IMF (Industrial manufacture); PREP (Preparation)
        (biphasic polymn. process using hydrolysis-sensitive monomers)
RE.CNT
RE
(1) Kochanowski; US 4286083 A 1981 HCAPLUS
(2) Rutgers The State University; WO 9630331 A 1996 HCAPLUS
L92
     ANSWER 15 OF 54 HCAPLUS COPYRIGHT 2001 ACS
AN
     1999:24346 HCAPLUS
DN
     130:182998
TΙ
     The study of water uptake in degradable polymers by thermally
     stimulated depolarization currents
     Suarez, Nery; Brocchini, Stephen; Kohn, Joachim
ΑU
CS
     Physics Department, Universidad Simon Bolivar, Caracas, Venez.
     Biomaterials (1998), 19(24), 2347-2356
CODEN: BIMADU; ISSN: 0142-9612
SO
PB
     Elsevier Science Ltd.
DΤ
     Journal
LΑ
     English
AΒ
     Poly(DTH succinate) is a new, degradable polymer with potential
     applications as a medical implant material. This polymer can be
     classified as an alternating copolymer of succinic acid and
     desaminotyrosyl-tyrosine hexyl ester (DTH), a diphenolic monomer
     derived from the natural amino acid L-tyrosine. In this study,
     the effects of water uptake (hydration) on the secondary relaxations of
     poly(DTH succinate) were investigated using the technique of thermally
     stimulated depolarization currents (TDSC). Four relaxation peaks were
     precisely characterized by means of Gaussian activation energy
     distributions. Drying and rehydration treatments show that only small
     amts. of water, at most 0.5% (wt./wt.), are necessary to hydrate two of
     the four polar moieties in poly(DTH succinate): the pendent chain ester
     carbonyls and the amide carbonyls in the polymer backbone.
     Water appeared to be more tightly bound to the amide carbonyl group and
     more loosely bound to the ester carbonyl group in the pendent chain. Even
     at a high state of hydration (1.2% wt./wt.), TDSC indicated that no water
     was assocd. with the Ph ester bond in the polymer backbone.
     This finding may explain the unexpectedly high stability of this
     polymer toward hydrolysis under physiol. conditions.
     Polymer packing was also affected by hydration. In an
     intermediate hydration state (water content: 0.5% wt./wt.) polymer
     packing was less dense than in the wet state (water content: 1.2%
     wt./wt.). This study represents the first application of the TDSC
     technique to the study of hydration in a degradable biomedical
     polymer. The results obtained indicate that the TDSC technique
     may be useful to assist in the understanding of the mechanisms of
     hydration and subsequent hydrolytic degrdn. in degradable biomaterials.
ΙT
     149787-38-2
     RL: PEP (Physical, engineering or chemical process); PRP (Properties);
     PROC (Process)
        (water uptake in degradable polymers studied by thermally
        stimulated depolarization currents)
     149787-38-2 HCAPLUS
RN
```

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

 ${\rm HO_2C-CH_2-CH_2-CO_2H}$ 

## IT 149787-38-2

RL: PEP (Physical, engineering or chemical process); PRP (Properties); PROC (Process)

(water uptake in degradable polymers studied by thermally

stimulated depolarization currents)

RE.CNT 35

RF.

- (1) Aldana, M; J Polym Sci Part B 1994, V32(13), P2197 HCAPLUS
- (2) Brocchini, S; J Amer Chem Soc 1997, V119(19), P4553 HCAPLUS
- (3) Cho, C; J Control Rel 1988, V7, P283 HCAPLUS
- (4) Colmenero, J; Makromol Chem (Macromol Symp) 1988, V20/21, P397 HCAPLUS
- (5) Crommen, J; Biomaterials 1992, V13(9), P601 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L92 ANSWER 16 OF 54 HCAPLUS COPYRIGHT 2001 ACS
- AN 1998:584001 HCAPLUS
- DN 129:306444
- TI Structure-property correlations in a combinatorial
  - library of degradable biomaterials
- AU Brocchini, Stephen; James, Kenneth;
  - Tangpasuthadol, Varawut; Kohn, Joachim
- CS Department of Chemistry, Rutgers, State University of New Jersey, New Brunswick, NJ, 08903, USA
- SO J. Biomed. Mater. Res. (1998), 42(1), 66-75
- CODEN: JBMRBG; ISSN: 0021-9304
- PB John Wiley & Sons, Inc.
- DT Journal
- LA English
- AB A combinatorial library of degradable polyarylates was prepd. These polymers are A-B-type copolymers consisting of an alternating sequence of a diphenol and a diacid. The library was prepd. by copolymg., in all possible combinations, 14 different tyrosine-derived diphenols and eight

different aliph. diacids, resulting in  $8 \times 14 = 112$  distinct polymers. This approach (a) increases the no. of available polymeric candidate materials for medical applications, and (b) facilitates the identification of correlations between polymer structure and glass transition temp., air-water contact angle, mech. properties, and fibroblast proliferation. The pendent chain and back-bone structures were systematically varied by (a) simple homologative variations in the no. of methylene groups, (b) substitution of oxygen for methylene groups, and (c) introduction of branched and arom. structures. The polymers contained within the library exhibited incremental variations in Tg (from 2.degree.C to 91.degree.C) and air-water contact angle (from 640 to 1010). Fibroblast proliferation (in vitro, serum-contg. media) ranged from approximating that measured on tissue culture polystyrene to complete absence of proliferation. Generally, decreased proliferation correlated linearly with increased surface hydrophobicity, except in those polymers derived from oxygen-contg. diacids in their backbone which were uniformly good growth substrates even if their surfaces were very hydrophobic. In a selected subgroup of polymers, tensile strength of thin solvent cast films ranged from about 6 to 45 MPa, while Young's modulus (stiffness) ranged from about 0.3 to 1.7 GPa. Combinatorial biomaterial libraries such as these tyrosine-derived polyarylates permit the systematic study of material-dependent biol. responses and provide the medical device designer with the option to choose a suitable material from a library of related polymers that encompasses a broad range of properties.

IT 149787-38-2P

RL: BAC (Biological activity or effector, except adverse); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(structure-property correlations in a combinatorial

library of degradable biomaterials)

149787-38-2 HCAPLUS

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

CM 1

RN CN

> CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

· HO2C-СH2-СH2-СО2Н

```
ΙT
     149787-38-2P 149787-41-7P 149826-02-8P
     188712-92-7P 188712-95-0P 188712-97-2P
     188712-99-4P 188713-01-1P 188713-03-3P
     188713-05-5P 188713-09-9P
     RL: BAC (Biological activity or effector, except adverse); PRP
     (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (structure-property correlations in a combinatorial
      library of degradable biomaterials)
L92
     ANSWER 17 OF 54 HCAPLUS COPYRIGHT 2001 ACS
AN
     1998:531630 HCAPLUS
DN
     129:293785
TΙ
     Biosmart tyrosine polycarbonates
ΑU
     Brode, G. L.; Kohn, J.; Kemnitzer, J. E.
CS
     Integra LifeSciences Corp., Plainsboro, NJ, 08536, USA
SO
     Polym. Prepr. (Am. Chem. Soc., Div. Polym. Chem.) (1998), 39(2), 230-231
     CODEN: ACPPAY; ISSN: 0032-3934
PB
     American Chemical Society, Division of Polymer Chemistry
DT
     Journal
LA
     English
     Poly(desaminotyrosyltyrosine-co-desaminotyrosyltyrosine carbonate) was
AΒ
     prepd. and was sol. at pH.gtoreq.6.0 in aq. or ionic buffer. Relevant
     medical applications are post-surgical antiadhesion barriers and oral drug
     delivery.
IT
     214259-59-3DP, hydrogenated
     RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Biosmart tyrosine polycarbonates)
     214259-59-3 HCAPLUS
RN
    L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, polymer with carbonic
CN
     acid and N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine ethyl ester (9CI)
     (CA INDEX NAME)
    CM
         1
    CRN 135313-59-6
    CMF C20 H23 N O5
    CDES 5:L
```

Absolute stereochemistry.

CM 2

CRN 86432-31-7 CMF C18 H19 N O5 CDES 5:L

Absolute stereochemistry.

CM 3

CRN 463-79-6 CMF C H2 O3

IT 214259-59-3DP, hydrogenated

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (Biosmart tyrosine polycarbonates)

L92 ANSWER 18 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1998:417621 HCAPLUS

DN 129:127136

TI Comparative histological evaluation of new tyrosine-derived polymers and poly(L-lactic acid) as a function of polymer degradation

AU Hooper, Kimberly A.; Macon, Natalie D.; Kohn, Joachim

CS Department of Chemistry, Rutgers-The State University of New Jersey, Piscataway, NJ, 08854-8087, USA

SO J. Biomed. Mater. Res. (1998), 41(3), 443-454 CODEN: JBMRBG; ISSN: 0021-9304

John Wiley & Sons, Inc.

PB John Wi DT Journal

LA English

AΒ

Previous studies demonstrated that poly(DTE carbonate) and poly(DTE adipate) (DTE = deaminotyrosyltyrosine Et ester) have suitable properties for use in biomedical applications. This study reports the evaluation of the in vivo tissue response to these polymers in comparison to poly(L-lactic acid) (PLLA). Typically, the biocompatibility of a material is detd. through histol. evaluations as a function of implantation time in a suitable animal model. However, due to changes that can occur in the tissue response at different stages of the degrdn. process, a fixed set of time points is not ideal for comparative evaluations of materials having different rates of degrdn. Therefore the tissue response elicited by poly(DTE carbonate), poly(DTE adipate), and PLLA was evaluated as a function of mol. wt. This allowed the tissue response to be compared at corresponding stages of degrdn. Poly(DTE adipate) consistently elicited the mildest tissue response, as judged by the width and lack of cellularity of the fibrous capsule formed around the implant. The tissue response to poly(DTE carbonate) was mild throughout the 570 day study. However, the response to PLLA fluctuation of the degree of degrdn., exhibiting an increase in the intensity of inflammation as the implant began to lose mass. At the completion of the study, tissue ingrowth into the degrading and disintegrating poly(DTE adipate) implant was evident while no comparative ingrowth of tissue was seen for PLLA. The similarity of the in vivo and in vitro degrdn. rates of each polymer confirmed the absence of enzymic involvement in the degrdn. process. A comparison of mol. wt. retention, water uptake, and mass loss in vivo with two commonly used in vitro systems [phosphate-buffered saline (PBS) and simulated body

fluid (SBF)] demonstrated that for the two tyrosine-derived polymers the in vivo results were equally well simulated in vitro with PBS and SBF. However, for PLLA the in vivo results were better simulated in vitro using PBS.

IT 149787-41-7

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (histol. evaluation of implants as function of polymer degrdn.)

RN 149787-41-7 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, ethyl ester, polymer with hexanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 135313-59-6 CMF C20 H23 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 124-04-9 CMF C6 H10 O4

 $HO_2C^-$  (CH<sub>2</sub>)<sub>4</sub> - CO<sub>2</sub>H

## IT 149787-41-7 174702-85-3

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (histol. evaluation of implants as function of polymer degrdn.)

L92 ANSWER 19 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1998:342334 HCAPLUS

DN 129:54839

TI High temperature signals for thermal-stimulated polarization and depolarization experiments in poly(DTH succinate)

AU Puma, M.; Suarez, N.

CS POBA International #438, Instituto de Ingenieria, Miami, FL, 33102-5255, USA

SO J. Appl. Polym. Sci. (1998), 69(2), 283-291 CODEN: JAPNAB; ISSN: 0021-8995

PB John Wiley & Sons, Inc.

DT Journal

LA English

AB Thermal-stimulated polarization and depolarization expts. with and without blocking electrodes are performed on a particular tyrosine-derived polyarylate: poly(DTH succinate). The high temp. region comprising the combined effects of the glass transition relaxation peak and the conduction through the sample is modeled. Conduction through the sample is described by a simple temp. relaxation model that, in the presence of blocking electrodes, gives rise to a charge redistribution peak. The anal. expression for this peak is found and, together with a convenient

description for the glass transition relaxation peak the exptl. data, is closely reproduced. An est. of the dielec. const. can be obtained with the model proposed. For the sample used, the value is equal to 2.4. 149787-38-2

RL: PEP (Physical, engineering or chemical process); PRP (Properties); PROC (Process)

(high temp. signals for thermal-stimulated polarization and depolarization expts. in poly(desaminotyrosyltyrosine hexyl succinate)) 149787-38-2 HCAPLUS

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

CM 1

ΙT

RN CN

> CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

 ${\tt HO_2C-CH_2-CH_2-CO_2H}$ 

## IT 149787-38-2

RL: PEP (Physical, engineering or chemical process); PRP (Properties); PROC (Process)

(bigh temp signals for thermal-stimulated polarization and

(high temp. signals for thermal-stimulated polarization and depolarization expts. in poly(desaminotyrosyltyrosine hexyl succinate))

L92 ANSWER 20 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:738720 HCAPLUS

DN 128:26780

TI Amino acid derived **polymers** for use in controlled delivery systems of peptides

AU Brocchini, S.; Schachter, D. M.; Kohn, J.

CS Department of Chemistry, Rutgers, The State University of New Jersey, New Brunswick, NJ, 08903, USA

SO ACS Symp. Ser. (1997), 675 (Therapeutic Protein and Peptide Formulation and Delivery), 154-167 CODEN: ACSMC8; ISSN: 0097-6156

PB American Chemical Society

DT Journal; General Review

LA English

AB A review with 32 refs. A family of synthetic, tyrosine-derived polyarylates is being studied as a new polymeric matrix system for the controlled release of peptides. The polyarylates are degradable amorphous materials whose backbone structure contains amide bonds. Using

the cyclic heptapeptide contained in the platelet integrin glycoprotein IIb/IIIa blocking formulation INTEGRILIN as a model, the effect of the polymer structure on peptide miscibility within the polymeric matrix and its effect on the release behavior was investigated. A new co-pptn. technique provided polyarylate-peptide blends that were compression molded without decompn., deactivation, or detectable aggregation of the peptide. Transparent, pliable compression molded films with high peptide loadings of up to 50% (wt./wt.) were fabricated in this way. In spite of such high loadings, the model peptide was not released from these films over a 30 day exposure to physiol. buffer soln. at 37 .degree.C. Only when polyethylene glycol (PEG) was added to the formulation was the model peptide released. Release rate was a function of polyarylate structure and the amt. and mol. wt. of PEG used in the blends. This provided an effective means to modulate the release rate.

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L92
     ANSWER 21 OF 54 HCAPLUS COPYRIGHT 2001 ACS
     1997:465129 HCAPLUS
ΑN
DN
     127:82246
ΤI
     Block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide)
     for biocompatible articles
IN
     Kohn, Joachim B.; Yu, Chun
     Rutgers, the State University, USA; Kohn, Joachim B.; Yu, Chun
PA
     PCT Int. Appl., 34 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
LΑ
FAN.CNT 3
                       KIND
     PATENT NO.
                             DATE
                                             APPLICATION NO.
     WO 9719996
                        A1
                             19970605
                                             WO 1996-US19098
                                                               19961127 <--
PΙ
         W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK,
             EE, ES, FI, GB, GE, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK,
             LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML,
             MR, NE, SN, TD, TG
     US 5658995
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                                             US 1995-562842
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                                                               19951127 <--
     CA 2237578
                             19970605
                                             CA 1996-2237578
                                                               19961127 <--
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     AU 9711264
                             19970619
                                             AU 1997-11264
                                                               19961127 <--
                        Α1
     AU 722764
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                        B2
     EP 863946
                        A1
                             19980916
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             IE. FI
     JP 2000501139
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                        Т2
                                                               19961127 <--
     US 6048521
                             20000411
                                             US 1998-85571
                        Α
                                                               19980527 <--
PRAI US 1995-562842
                       19951127
                                 <--
     WO 1996-US19098
                      19961127
                                 <--
     US 1997-64656
                       19971107
                                 <--
     US 1997-64905
                       19971107
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     US 1998-81502
                       19980413
                                <--
AB
     Random block copolymers [0-p-C6H4R1CONHCH(CO2R2)CH2C6H4-p-OCO]1-
     f[(OR3)yOCO]f[I; R1 = CH:CH or (CH2)j, j = 0 or 1-8; R2 = H, straight and
     branched alkyl and alkylaryl groups contg. .ltoreq.18 C atoms and their
     derivs. or biol. and pharmaceutically active compds. covalently bonded to
     the copolymer; R3 = alkylene group contg. .ltoreq.4 C atoms; y =
     .apprx.5-3000; and f = .apprx.1-99 \text{ mol} and polyarylate block copolymers
     are prepd. by the polymn. of tyrosine-based diphenol, dicarboxylic acid
     (deriv.)., and a polyalkylene oxide. The block copolymers are useful for
     implantable medical devices and drug delivery implants. The
     desaminotyrosyl tyrosine Et ester-phosgene-polyethylene glycol (5%) block
     copolymer of no.-av. mol. wt. 84,000 daltons had an initial tensile
     strength 1.3 GPa.
ΙT
     191858-74-9
```

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(lblock copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

RN 191858-74-9 HCAPLUS

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride and .alpha.-hydro-.omega.-hydroxypoly(oxy-1,2-ethanediyl), block (9CI) (CA INDEX NAME)

CM 1

CN

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 25322-68-3 CMF (C2 H4 O)n H2 O CCI PMS

$$HO \longrightarrow CH_2 - CH_2 - O \longrightarrow n$$

CM 3

CRN 75-44-5 CMF C C12 O

IT 191858-74-9

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(lblock copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

IT 191858-68-1P 191858-70-5P

RL: IMF (Industrial manufacture); PRP (Properties); THU (Therapeutic use);
BIOL (Biological study); PREP (Preparation); USES (Uses)
 (block copolymers of tyrosine-based polycarbonate and poly(alkylene oxide) for biocompatible articles)

IT 191858-72-7

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES

(block copolymers of tyrosine-based polycarbonate and poly(alkylene

oxide) for biocompatible articles) ANSWER 22 OF 54 HCAPLUS COPYRIGHT 2001 ACS L92 AN 1997:342358 HCAPLUS DN 126:330869 ΤI A Combinatorial Approach for Polymer Design ΑU Brocchini, Stephen; James, Kenneth; Tangpasuthadol, Varawut; Kohn, Joachim CS Department of Chemistry, Rutgers The State University of New Jersey, New Brunswick, NJ, 09803, USA J. Am. Chem. Soc. (1997), 119(19), 4553-4554 SO CODEN: JACSAT; ISSN: 0002-7863 PB American Chemical Society DTJournal English LA The design of specialty polymers for advanced applications is AΒ challenging since such polymers must meet stringent performance requirements while also exhibiting appropriate physicomech. properties suitable for a given application. The authors report the concept of creating a permutationally designed library of strictly alternating A-B type copolymers in which a set of monomers A provides a reactive group for the attachment of pendent chains and a set of monomers B provides a means for the structural variation of the polymer backbone. The copolymn. of each of the monomers in set A with each of the monomers in set B creates a "two-dimensional array" of polymers that can be used (a) to increase the no. of available polymer candidate materials for a given application, and (b) to systematize the study of structure-property correlations. approach may be particularly useful for the design of biodegradable polymers for medical applications: As a first implementation of this combinatorial approach, a library of 112 biodegradable polyacrylates was prepd. from 14 tyrosine-derived diphenols and 8 aliph. diacids. The 112 polymers were prepd. on a 0.2 g scale in arrays of glass vial set up in a shaker bath. The polyarylates were characterized by mol. wt., Tg, and air-water contact angle. Tg values ranged from 2-91 .degree.C and contact angles ranged from 64-101.degree. in a continuous incremental fashion and several structure-property correlations were obsd. In biomaterials research, it is of central importance to understand the influence of polymer structure on the cellular response. A seven day cellular proliferation study using rat lung fibroblasts established correlations between polyarylate structure and in vitro cell proliferation. ΙT 189760-07-4P RL: BAC (Biological activity or effector, except adverse); PRP (Properties); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation) (prepn. and cell proliferation properties of homologous series of polyarylates) 189760-07-4 HCAPLUS RNL-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, 2-(2-ethoxyethoxy)ethyl CN ester, polymer with 2,2'-oxybis[acetic acid] (9CI) (CA INDEX NAME) CM 1 CRN 189760-06-3 CMF C24 H31 N O7

Absolute stereochemistry.

(Uses)

CM 2

CRN 110-99-6 CMF C4 H6 O5

HO2C-CH2-O-CH2-CO2H

IT 189760-07-4P 189760-09-6P 189760-11-0P 189760-12-1P 189760-14-3P 189760-16-5P

RL: BAC (Biological activity or effector, except adverse); PRP (Properties); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(prepn. and cell proliferation properties of homologous series of polyarylates)

L92 ANSWER 23 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:336802 HCAPLUS

DN 127:8987

TI Pseudo-poly(amino acid)s. Examples for synthetic materials derived from natural metabolites

AU James, Kenneth; Kohn, Joachim

CS Department of Chemistry, Rutgers - The State University of New Jersey, New Brunswick, NJ, 08903, USA

SO Controlled Drug Delivery (1997), 389-403. Editor(s): Park, Kinam. Publisher: American Chemical Society, Washington, D. C. CODEN: 64KFAX

DT Conference; General Review

LA English

A review with 57 refs. Pseudo-poly(amino acid)s consisting of alpha.-L-amino acid building blocks linked by nonamide bonds have been used successfully as carriers in direct intracranial drug delivery, immunizations systems, and the controlled release of anticoagulants. These natural metabolite-based polymers are biocompatible, degradable materials readily processed into microspheres, films, fibers, pins, and screws. In the last 5 yr, most attention has been directed toward tyrosine-derived polycarbonates, polyiminocarbonates, and polyarylates. Pseudo-poly(aminoacid) chem. and synthesis, physicomech. and degrdn. properties, biol. response, immunol. considerations, sterilization, and drug delivery applications thus far investigated are summarized.

L92 ANSWER 24 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:230323 HCAPLUS

TI A combinatorial approach for the development of new polymer biomaterials.

AU Brocchini, S.; James, K. S.; Tangpasuthadol, V.; Kohn, J.

CS Department Chemistry, Rutgers University, Piscataway, NJ, 08855, USA

SO Book of Abstracts, 213th ACS National Meeting, San Francisco, April 13-17 (1997), BIOC-257 Publisher: American Chemical Society, Washington, D. C. CODEN: 64AOAA

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DT
     Conference; Meeting Abstract
LA
     Development of synthetic degradable polymers for use in medical
ΑB
     applications including tissue replacement therapy is a complex challenge.
     These medical polymers must meet a multitude of stringent
     requirements similar in kind to drug candidates while also exhibiting
     appropriate physicomech. properties that meet the criteria for a given
     application. Of central importance is the requirement to understand the
     influence of polymer structure on the cellular response. We
     report the concept of creating a permutationally designed polymer
     system to prep. a library of polymers that can be used
     (a) to increase the no. of available polymer candidate materials
     for medical applications, and (b) to systematize the study of correlations
    between polymer structure, material properties, and biol.
     responses. In this combinatorial approach, structural
     variations of a small no. of monomers were systematically translated into
     a library of 112 tyrosinederived polyarylates which
     were prepd. on a 0.2 g scale in arrays of glass vials set up in a shaker
     bath. Up to 32 simultaneous reactions were conducted in a run and the
     polyarylates were characterized by mol. wt., Tg, and air-water contact
     angle. Properties, including in vitro fibroblast proliferation,
     incrementally varied over a wide range. This resulted in several
     structure-property correlations which will be described.
L92 ANSWER 25 OF 54 HCAPLUS COPYRIGHT 2001 ACS
AN
     1997:226789 HCAPLUS
DN
     126:216672
TΙ
     Polymeric drug formulations
    Brocchini, Stephen; Hanson, Stephen R.; Kohn, Joachim B.
IN
     Rutgers, the State University;, USA; Emory University
PΑ
SO
     PCT Int. Appl., 52 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                     KIND DATE
                                          APPLICATION NO. DATE
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                                           _____
                                          WO 1996-US11766 19960716 <--
                           19970213
     WO 9704744
                     A1
           AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE,
             ES, FI, GB, GE, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS,
            LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD,
             SE, SG
         RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR,
             IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN
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     JP 11510160
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                            19990907
PRAI US 1995-508577
                      19950728 <--
     WO 1996-US11766 19960716 <--
AB
     of a water-sol. drug in a water-insol. tissue-compatible polymer matrix.
     Polymeric drug formulations are also disclosed contg. a single-phase
     dispersion of a water-sol. drug and a water-insol. tissue-compatible
    polymer matrix, and a second, phase-disrupting polymer that is
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Polymeric drug formulations contg. a non-releasing single-phase dispersion of a water-sol. drug in a water-insol. tissue-compatible polymer matrix. Polymeric drug formulations are also disclosed contg. a single-phase dispersion of a water-sol. drug and a water-insol. tissue-compatible polymer matrix, and a second, phase-disrupting polymer that is non-miscible with the tissue-compatible polymer and is present in an amt. sufficient to form phase-sepd. microdomains of the second polymer in the tissue-compatible polymer matrix, so that the release rate of the water-sol. drug from the tissue-compatible polymer matrix is related to the amt. of the second polymer. Methods of prepg. the polymeric drug formulations are also described, as well as methods for site-specific drug delivery utilizing the polymeric drug formulations. Examples of

platelet-aggregation-inhibiting peptide formulations are given.

IT 149787-39-3

RL: PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (polymeric drug formulations)

RN 149787-39-3 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with hexanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 124-04-9 CMF C6 H10 O4

## HO2C-(CH2)4-CO2H-

## IT 149787-39-3

RL: PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (polymeric drug formulations)

L92 ANSWER 26 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:155759 HCAPLUS

DN 126:255433

TI Comparison of the effect of ethylene oxide and .gamma.-irradiation on selected tyrosine-derived polycarbonates and poly(L-lactic acid)

AU Hooper, Kimberly A.; Cox, J. David; Kohn, Joachim

CS Dep. Chem., Rutgers, State Univ. New Jersey, New Brunswick, NJ, 08903, USA

SO J. Appl. Polym. Sci. (1997), 63(11), 1499-1510

CODEN: JAPNAB; ISSN: 0021-8995

PB Wiley

DT Journal

LA English

Tyrosine-derived polycarbonates are a new class of degradable polymers that have possible biomedical applications. In this study, the effect of the two most common sterilization techniques, ethylene oxide and .gamma.-irradn. (0.3, 1.1, 3.9, 6.4, 10.6 Mrad), was evaluated for a family of four structurally related tyrosine-derived polycarbonates and for poly(L-lactic acid) (PLLA). The four polycarbonates were poly(DTE carbonate), poly(DTB carbonate), poly(DTB carbonate), and poly(DTO carbonate) and differed only in the length of the pendent chain. Ethylene oxide exposure had little effect on mol. wt., surface compn., mech. properties, or degran. rate of all test polymers except for poly(DTO

carboante). Poly(DTO carboante) was unique since following ethylene oxide exposure it degraded faster than did the nonsterilized control. .gamma.-Irradiated tyrosine-derived polycarbonates retained over 81% of their initial mol. wt. when exposed to a clin. relevant dose of 3.9 Mrad and retained still 58% of the initial mol. wt. when exposed to the highest test dose of 10.6 Mrad. No changes in surface compn. and only slight changes in yield strength and the Young's modulus were detected for any of the tyrosine-derived polycarbonates following .gamma.-irradn. In vitro, irradiated films of poly(DTE carbonate), poly(DTB carbonate), and poly(DTH carbonate) degraded at approx. the same rate as did the nonsterilized films regardless of irradn. dose. Only poly(DTO carbonate), irradiated at high doses, degraded faster than did the control. Medical-grade PLLA was tested under identical conditions. Ethylene oxide exposure of PLLA did not affect the mol. wt., surface compn., mech. properties, or in vitro degrdn. rate. However, upon irradn. at 10.6 Mrad, PLLA retained only 29% of its initial mol. wt.; a dose of 3.9 Mrad resulted in retention of 49% of the initial mol. wt. In correspondence with earlier publications, irradn. of PLLA induced significant losses in the Young's modulus, % strain at break, and changes in the postirradn. rate of degrdn. in some specimens. Compared to PLLA, tyrosine-derived polycarbonates are significantly more stable to .gamma.-irradn. and can be sterilized by conventional .gamma.-sterilization techniques.

IT 183480-53-7

RL: PRP (Properties); RCT (Reactant); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(ethylene oxide and .gamma.-irradn. effect on tyrosine-derived polycarbonates and poly(lactic acid))

RN 183480-53-7 HCAPLUS

CN Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-(butoxycarbonyl)-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

### IT 183480-53-7 183480-55-9

RL: PRP (Properties); RCT (Reactant); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(ethylene oxide and .gamma.-irradn. effect on tyrosine-derived polycarbonates and poly(lactic acid))

L92 ANSWER 27 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:155718 HCAPLUS

DN 126:255430

TI Surface characterization of tyrosine-derived polycarbonates

AU Perez-Luna, Victor H.; Hooper, Kimberly A.; Kohn, Joachim; Ratner, Buddy D.

CS Dep. Chem. Eng., Univ. Washington, Seattle, WA, 98195, USA

SO J. Appl. Polym. Sci. (1997), 63(11), 1467-1479 CODEN: JAPNAB; ISSN: 0021-8995

PB Wiley

DT Journal

LA English

The surfaces of five biodegradable tyrosine-derived polycarbonates were studied using contact angle measurements, ESCA, and static SIMS. The wettability, crit. surface tension, and polarity of these polymers decreased with increasing chain length of the pendent alkyl groups. Surface elemental compn., as detd. by ESCA, was consistent with the

stoichiometry of the repeat unit of the polymers. High-resoln. Cls, Ols, and Nls ESCA spectra also showed results consistent with the different bonding states of these elements in the polymer repeat unit. In both pos. and neg. ion spectra, SIMS expts. showed fragment ions characteristic of the polymer backbone. Fragment ions characteristic of the pendent groups were identified in the neg. ion SIMS spectra only, while the pos. SIMS spectra provided a characteristic fingerprint for each polymer.

IT 183480-53-7

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(surface characterization of tyrosine-derived polycarbonates)

RN 183480-53-7 HCAPLUS

CN Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-(butoxycarbonyl)-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

# IT 183480-53-7 183480-54-8 183480-55-9 188716-30-5

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(surface characterization of tyrosine-derived polycarbonates)

L92 ANSWER 28 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:155713 HCAPLUS

DN 126:255428

TI Molecular relaxation mechanisms of tyrosine-derived polycarbonates by thermally stimulated depolarization currents

AU Suarez, N.; Laredo, E.; Bello, A.; Kohn, J.

CS Physics Dep., Universidad Simon Bolivar, Caracas, Venez.

SO J. Appl. Polym. Sci. (1997), 63(11), 1457-1466

CODEN: JAPNAB; ISSN: 0021-8995

PB Wiley

DT Journal

LA English

AB A series of tyrosine-derived polycarbonates with different lengths (2 .ltoreq. n .ltoreq. 8) for the alkyl ester pendent chain were studied by measuring thermally stimulated depolarization currents (TSDC). The obsd. spectra could be sepd. into three regions: the low-temp. zone with a broad, complex .beta. band (80-240 K), the intermediate zone (250-300 K), and the high-temp. zone (300-400 K) with a sharp .alpha. peak. The application of direct signal anal. (DSA) to decomp. the complex peaks into elementary processes led to the detn. of the relaxation time distribution and temp. dependence of each process. The variation of the relaxation parameters as a function of the pendent chain length facilitated the tentative identification of the relaxation mechanisms responsible for the obsd. current peaks. It is proposed that as the temp. increases one observes, first, the individual motion of each polar group, then the concerted motion of the entire pendent chain, and, last, the movement of the polymer backbone.

IT 183480-53-7

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(mol. relaxation mechanisms of tyrosine-derived polycarbonates by thermally stimulated depolarization currents)

RN 183480-53-7 HCAPLUS

Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-(butoxycarbonyl)-1,2-CN ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

183480-53-7 183480-54-8 183480-55-9 IT

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES

(mol. relaxation mechanisms of tyrosine-derived polycarbonates by thermally stimulated depolarization currents)

L92 ANSWER 29 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN1997:155655 HCAPLUS

DN 126:255427

TI Thermal properties and enthalpy relaxation of tyrosine-derived polyarylates

Tangpasuthadol, Varawut; Shefer, Adi; Yu, Chun; Zhou, Jing; ΑU Kohn, Joachin

Dep. Chem., Rutgers, State Univ. New Jersey, New Brunswick, NJ, 08903, USA CS

SO J. Appl. Polym. Sci. (1997), 63(11), 1441-1448

CODEN: JAPNAB; ISSN: 0021-8995

PΒ Wiley

AB

DT Journal

English LA

Sixteen degradable, tyrosine-derived polyarylates with well defined chem. structures were used to study the effect of polymer structure on the glass transition temp. and enthalpy relaxation kinetics (phys. aging). These polyarylates compose a model system where the no. of methylene groups present in either the pendent chain or the polymer backbone can be altered independently and in a systematic fashion. Quant. differential scanning calorimetry was employed to measure the glass transition temp. and the enthalpy relaxation kinetics. Correlations between these material properties and the polymer structure were established. The glass transition temp. of this family of polymers ranged from 13 to 78.degree.C. The addn. of methylene groups to either the pendent chain or the polymer backbone made a fairly const. contribution to lowering the glass transition the glass transition temp. The rate of enthalpy relaxation increased with an increasing no. of methylene groups in the polymer backbone, but was independent of the no. methylene groups in the pendent chain. observation indicated that the rate of enthalpy relaxation in these polymers was limited by the mobility of the polymer backbone. The enthalpy relaxation data was fitted to the Cowie-Ferguson model and the relaxation times obtained ranged from 44 min to about 100 min. Although these structure-property correlations facilitate the design of new materials with predictable thermal properties, they are rarely investigated for biomedical polymers. 149787-38-2

ΙT

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(thermal properties and enthalpy relaxation of tyrosine -derived polyarylates)

149787-38-2 HCAPLUS RN

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer CN with butanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

HO2C-СH2-СH2-СО2Н

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IT 149787-38-2 149787-39-3 149787-40-6
149787-41-7 149826-02-8 188712-92-7
188712-95-0 188712-97-2 188712-99-4
188713-01-1 188713-03-3 188713-05-5
188713-08-8 188713-09-9 188713-10-2
188713-11-3
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RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(thermal properties and enthalpy relaxation of **tyrosine** -derived polyarylates)

L92 ANSWER 30 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:36235 HCAPLUS

DN 126:65248

TI New biomaterials for tissue engineering

AU James, Kenneth; Kohn, Joachim

CS USA

SO MRS Bull. (1996), 21(11), 22-26 CODEN: MRSBEA; ISSN: 0883-7694

PB Materials Research Society

DT Journal; General Review

LA English

AB A review with 41 refs., esp. on tyrosine-derived polycarbonates and polyarylates.

L92 ANSWER 31 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:14265 HCAPLUS

TI Applications of pseudo-poly(amino acid) biomaterials

AU James, Kenneth; Kohn, Joachim

CS Dep. Chem., Rutgers Univ., Piscataway, NJ, 08855-0939, USA SO Trends Polym. Sci. (Cambridge, U. K.) (1996), 4(12), 394-397

CODEN: TPSCE8; ISSN: 0966-4793

PB Elsevier

DT Journal: News Announcement

LA English

AB Unavailable

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ANSWER 32 OF 54 HCAPLUS COPYRIGHT 2001 ACS
L92
     1996:710485 HCAPLUS
AN
     126:8702
DN
     Improved synthesis of tyrosine-derived diphenol monomers
ΤI
     Kohn, Joachim B.; Hooper, Kimberly Ann; Brocchini, Stephen
IN
     J.; Schwartz, Arthur L.
     Rutgers, the State University, USA
PA
     PCT Int. Appl., 26 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
LA
FAN.CNT 1
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                                                             DATE
     PATENT NO.
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                            19961003
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                       A1
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             SG, SI
         RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR,
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     JP 11503421
                      19950331 <--
PRAI US 1995-414339
                      19960329
                                <--
     US 1996-625763
                      19960329 <--
     WO 1996-US4387
     MARPAT 126:8702
os
GΙ
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HO R1CONHCHCH<sub>2</sub> OH 
$$CO_2R^2$$

A method for prepg. diphenol compds. [I; R1 = CH:CH, (CH2)n; wherein n =AB 0, 1-8; R2 = linear or branched C.ltoreq.18 alkyl or aralkyl] involves the steps of coupling a hydroxyphenyl carboxylic acid HOC6H4-R1-CO2H (R1 = same as above) with an L-tyrosine ester HOC6H4-CH2CH(CO2R2)NH2 (R2 = same as above) in a water-miscible org. reaction solvent contg. a carbodiimide capable of forming a water-sol. urea byproduct, thereby forming a diphenol reaction product and combining the reaction mixt. with an amt. of water effective to ppt. the diphenol as a water-immiscible org. phase, so that a water-immiscible org. phase is formed contg. the diphenol reaction product. New diphenol monomers and polymers polymd. therefrom are also disclosed. Thus, 9.63 g hexyl Ltyrosinate was condensed with 6.04 g desaminotyrosine using 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide hydrochloride in THF in an ice bath for 1 h and at room temp. for 19 h to give 71% hexyl N-( desaminotyrosinyl)-L-tyrosinate of 95.8% purity. This diphenol was polymd. in soln. with phosgene to give poly[hexyl N-(desaminotyrosinyl)-L-tyrosinate carbonate]. ΙT 133418-81-2P

RL: SPN (Synthetic preparation); PREP (Preparation) (improved prepn. of alkyl desaminotyrosinyltyrosinate as

diphenol monomers for polycarbonate polymers)

RN 133418-81-2 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5

CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C C12 O

IT 133418-81-2P 183480-48-0P 183480-50-4P 183480-51-5P 183480-53-7P 183480-54-8P 183480-55-9P

RL: SPN (Synthetic preparation); PREP (Preparation) (improved prepn. of alkyl desaminotyrosinyltyrosinate as diphenol monomers for polycarbonate polymers)

L92 ANSWER 33 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1996:250260 HCAPLUS

DN 124:325298

TI Canine bone response to tyrosine-derived polycarbonates

and poly(L-lactic acid)

AU Choueka, Jack; Charvet, Jose L.; Koval, Kenneth J.; Alexander, Harold; James, Kenneth S.; Hooper, Kimberly A.; Kohn, Joachim

CS Dep. Bioengineering, Orthopaedic Inst., New York, NY, USA

SO J. Biomed. Mater. Res. (1996), 31(1), 35-41

CODEN: JBMRBG; ISSN: 0021-9304

DT Journal

LA English

AB Tyrosine-derived polycarbonates are a new class of degradable polymers developed for orthopedic applications. In this study the long-term (48 wk) in vivo degrdn. kinetics and host bone response to poly(DTE carbonate) and poly(DTH carbonate) were investigated using a canine bone chamber model. Poly(L-lactic acid) (PLA) served as a control material. Two chambers of each test material were retrieved at 6-, 12-, 24-, and 48-wk time points. Tyrosine-derived polycarbonates were found to exhibit degrdn. kinetics comparable to PLA. Each test material lost approx. 50% of its initial mol. wt. (Mw) over the 48-wk test period. Poly(DTE carbonate) and poly(DTH carbonate)

test chambers were characterized by sustained bone ingrowth throughout the 48 wk. In contrast, bone ingrowth into the PLA chambers peaked at 24 wk and dropped by half at the 48-wk time point. A fibrous tissue layer was found surrounding the PLA implants at all time points. This fibrous tissue layer was notably absent at the interface between bone and the tyrosine-derived polycarbonates. Histol. sections revealed intimate contact between bone and tyrosine-derived polycarbonates. From a degrdn.-biocompatibility perspective, the tyrosine-derived polycarbonates appear to be comparable, if not superior, to PLA in this canine bone chamber model.

L92 ANSWER 34 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1996:111206 HCAPLUS

DN 124:203129

TI Diphenolic monomers derived from the natural amino acid .alpha.-L-tyrosine: an evaluation of peptide coupling techniques

AU Hooper, Kimberly A.; Kohn, Joachim

CS Department of Chemistry, Rutgers, State University of New Jersey, New Brunswick, NJ, 08903, USA

SO J. Bioact. Compat. Polym. (1995), Volume Date 1995, 10(4), 327-40 CODEN: JBCPEV; ISSN: 0883-9115

DT Journal

LA English

AB Desaminotyrosyl-tyrosine Et, Bu, hexyl, and octyl esters were prepd. from tyrosine esters and desaminotyrosine derivs. The esters were diphenols and polycarbonates were prepd. from them.

IT 143715-03-1P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of diphenolic desaminotyrosyl-tyrosine esters and their polycarbonates)

RN 143715-03-1 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 463-79-6 CMF C H2 O3

#### 143715-03-1P 174702-85-3P 174702-86-4P ΙT 174702-87-5P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of diphenolic desaminotyrosyl-tyrosine esters and their polycarbonates)

ANSWER 35 OF 54 HCAPLUS COPYRIGHT 2001 ACS L92

1996:104965 HCAPLUS AN

124:211936 DN

Thermal properties and physical aging behavior of tyrosine ΤI -derived polycarbonates

Tangpasuthadol, Varawut; Shefer, Adi; Hooper, Kimberly A.; ΑU Kohn, Joachim

Dep. Chemistry, State Univ. New Jersey, New Brunswick, NJ, 08903, USA CS

Biomaterials (1996), 17(4), 463-8 SO

CODEN: BIMADU; ISSN: 0142-9612

DT Journal

LA English

AB Tyrosine-derived polycarbonates are new

carbonate-amide copolymers. These materials have been suggested for use in medical applications, but their thermal properties and their enthalpy relaxation kinetics (phys. aging behavior) have so far not been evaluated in detail. Since structure-property correlations involving enthalpy relaxation are rarely investigated for biomedical

polymers, a series of four tyrosine-derived

polycarbonates was used as a model system to study the effect of pendant chain length on the thermal properties and the enthalpy relaxation kinetics. The chem. structure of the test polymers was identical except for the length of their resp. pendant chains. This feature facilitated the identification of structure-property correlations. Quant. differential scanning calorimetry was utilized to det. the thermal properties and to measure enthalpy relaxation kinetics. The glass transition temp. of this family of polymers decreased from 93 to 52.degree.C when the length of the pendant chain was increased from two to eight carbon atoms. Successive addns. of methylene groups to the pendant chain made a fairly const. contribution to lowering the glass transition temp. For pendant chains of four or more methylene groups, the rate of enthalpy relaxation was independent of the no. of methylene groups in the pendant chain. The enthalpy relaxation data were fitted to the Cowie-Ferguson model and the relaxation times obtained were about 90 min. Dynamic mech. anal. was employed to study the viscoelastic properties. The available observations indicate that the polymers become more flexible with increasing length of the pendant chain. The results suggest that the length of the pendant chain can be used effectively to control important material properties in this series of polymers

#### ΙT 171436-75-2P

CN

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (thermal properties and phys. aging behavior of tyrosine

-derived polycarbonates)

RN171436-75-2 HCAPLUS

Poly[oxycarbonyloxy-1,4-phenylene[2-(ethoxycarbonyl)-1,2ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

# IT 171436-75-2P 171436-76-3P 171436-77-4P 171436-78-5P

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (thermal properties and phys. aging behavior of tyrosine -derived polycarbonates)

L92 ANSWER 36 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1995:927839 HCAPLUS

DN 124:15445

TI Evaluation of thermal properties and physical aging as function of the pendent chain length in tyrosine-derived polycarbonates , a class of new biomaterials

AU Tangpasuthadol, Varawut; Shefer, Adi; Hooper, Kimberly A.; Kohn, Joachim

CS Dep. Chem., Rutgers Univ., Piscataway, NJ, 08855-0939, USA

SO Mater. Res. Soc. Symp. Proc. (1995), 394 (Polymers in Medicine and Pharmacy), 143-8
CODEN: MRSPDH; ISSN: 0272-9172

DT Journal

LA English

Tyrosine-derived polycarbonates are currently in preclin. evaluations for biomedical applications. Although phys. aging can significantly alter a wide range of polymer properties, phys. aging is has rarely been investigated for biomedical polymers. A series of four tyrosine-derived polycarbonates was used as a model system to study the effect of polymer structure on the enthalpy relaxation kinetics.

IT 171436-75-2

171436-75-2
RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(thermal properties and phys. aging as function of pendent chain length in tyrosine-derived polycarbonates for biomaterials)

RN 171436-75-2 HCAPLUS

CN—Poly[oxycarbonyloxy=1,4=phenylene[2-(ethoxycarbonyl)-1,2ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

# IT 171436-75-2 171436-76-3 171436-77-4 171436-78-5

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(thermal properties and phys. aging as function of pendent chain length in tyrosine-derived polycarbonates for biomaterials)

- L92 ANSWER 37 OF 54 HCAPLUS COPYRIGHT 2001 ACS
- AN 1995:843546 HCAPLUS
- DN 124:37506
- TI The effect of polymer structure and formulation techniques on the release of a model peptide
- AU Kohn, J.; Brocchini, S.; Imai, M.; Vyavahare, N.
- CS Department Chemistry, Rutgers University, Piscataway, NJ, 08855, USA
- SO Proc. Int. Symp. Controlled Release Bioact. Mater. (1995), 22nd,

522-3

CODEN: PCRMEY; ISSN: 1022-0178

DT Journal

LA English

AB The availability of polymers that can be molded at temps. <100.degree. makes it possible to use compression molding, extrusion, or injection molding for the prepn. of controlled release systems contg. heat sensitive peptide drugs. In such systems, there is a need for the intimate, uniform dispersion of the drug withing the polymer matrix. This was accomplished with a copptn. technique. Using Integrelin as a model drug, the prolonged release of this peptide from thin compression molded films was demonstrated.

IT 149787-39-3

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(polymer structure and formulation techniques effect on release of a model peptide)

RN 149787-39-3 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with hexanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 124-04-9 CMF C6 H10 O4

 $HO_2C^-$  (CH<sub>2</sub>)<sub>4</sub> - CO<sub>2</sub>H

## IT 149787-39-3 149787-41-7 149826-02-8

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(polymer structure and formulation techniques effect on release of a model peptide)

L92 ANSWER 38 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1994:563954 HCAPLUS

DN 121:163954

TI Design, synthesis, and preliminary characterization of tyrosine-containing polyarylates: new biomaterials for medical applications

AU Fiordeliso, James; Bron, Samuel; Kohn, Joachim

CS Dep. Chem., Rutgers State Univ., New Brunswick, NJ, 09803, USA

SO J. Biomater. Sci., Polym. Ed. (1994), 5(6), 497-510 CODEN: JBSEEA; ISSN: 0920-5063

DT Journal

LA English

Five structurally related, aliph. polyarylates were prepd. from AB tyrosine-derived diphenols and diacids. The diphenols were a homologous series of 3 desaminotyrosyl-tyrosine alkyl esters (Et, hexyl, octyl) which had previously been used in the synthesis of mech. strong and tissue-compatible polycarbonates. The diacids (succinic acid, adipic acid, sebacic acid) were selected among compds. that were known to be of low systemic toxicity. By using different diacids as comonomers, the flexibility of the polymer backbone could be varied, while the desaminotyrosyl-tyrosine alkyl esters provided pendent chains of various length. Some of the thermal and mech. properties of the 5 polymers were correlated to their chem. structure: the glass transition temp. decreased from 53 to 13.degree., and the tensile modulus (measured at room temp.) decreased from 1500 to about 3 MPa when the length of the aliph. diacid in the polymer backbone and/or the length of the alkyl ester pendent chain was increased. The presence of an arylate bond in the polymer backbone introduced a hydrolytically labile linkage into the polymer structure. Under physiol. conditions in vitro all polymers degraded: thin films retained only about 30-40% of their initial mol. wt. (Mw) after 26 wk of storage in phosphate buffer solns. (pH 7.4) at 37.degree.. Release studies with p-nitroaniline as a model drug indicated that a diffusion controlled release process occurred. The rate of p-nitroaniline release was correlated with the glass transition temp. of the polymer.

IT 149787-38-2P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
 (prepn. and characterization of, as biomaterial for medical
 applications)

RN 149787-38-2 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

 $HO_2C-CH_2-CH_2-CO_2H$ 

IT 149787-38-2P 149787-39-3P 149787-40-6P 149787-41-7P 149826-02-8P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and characterization of, as biomaterial for medical

## applications)

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1.92
     ANSWER 39 OF 54 HCAPLUS COPYRIGHT 2001 ACS
ΑN
     1993:546603 HCAPLUS
DN
     119:146603
TI
     Polyarylate containing derivatives of the natural amino acid L-tyrosine
ΙN
     Kohn, Joachim B.; Fiordeliso, James J.
PA
     Rutgers, State Univ., USA
     U.S., 13 pp. Cont.-in-part of U.S. Ser. No. 804,767.
SO
     CODEN: USXXAM
DT
     Patent
LA
     English
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                      KIND DATE
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             SE, SK, UA, VN
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                      19920813 <--
     WO 1993-US7676
                      19930813 <--
AΒ
     Amino acid-derived diphenols are copolymd. with dicarboxylic acid by a
     carbodiimide-mediated direct polycondensation to form nontoxic bioerodible
     polyarylates for matrixes in controlled drug delivery systems and for
     biocompatible molded articles. For example, poly(desaminotyrosyl-tyrosine
     hexyl ester adipate) was prepd. and its thermal properties and degrdn.
     rates in physiol. conditions were studied.
IT
     149787-38-2P
     RL: PREP (Preparation)
        (prepn. of, for matrix in controlled drug delivery systems and for
        medical goods)
     149787-38-2 HCAPLUS
RN
CN
     L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer
     with butanedioic acid (9CI) (CA INDEX NAME)
     CM
          1
     CRN 133063-33-9
     CMF C24 H31 N O5
     CDES 5:L
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Absolute stereochemistry.

2 CM

CRN 110-15-6 CMF C4 H6 O4

HO2C-CH2-CH2-CO2H

#### IT 149787-38-2P 149787-39-3P 149787-40-6P 149787-41-7P 149787-42-8P 149826-02-8P

RL: PREP (Preparation)

(prepn. of, for matrix in controlled drug delivery systems and for medical goods)

L92 ANSWER 40 OF 54 HCAPLUS COPYRIGHT 2001 ACS

ΑN 1993:450276 HCAPLUS

DN 119:50276

Physicomechanical properties of biodegradable polymers for medical ΤI applications

ΑU Engelberg, I.; Kohn, J.

RAFAEL - Armament Dev. Author., Haifa, 31021, Israel CS

Isr. Mater. Eng. Conf., 5th (1991) 277-91 SO

CODEN: 58SIAV

DT Journal

English LA

The physicomech. properties of degradable polymers used for medical AΒ applications were characterized. The following polymers were included in this study: three samples of poly(ortho esters) derived from 3,9-bis(ethylidene-2,4,8,10-tetraoxaspiro[5.5]undecane) and various ratios of 1,6-hexanediol and trans-cyclohexanedimethanol, poly(glycolic acid), six samples of poly(L-lactic acid) and poly(D,L-lactic acid) with mol. wts. 21,000-550,000 dalton, poly(.epsilon.-caprolactone), poly(.beta.-hydroxybutyrate) and three copolymers of .beta.-hydroxybutyric acid and various amts. of hydroxyvaleric acid, one sample each of two different types of poly(anhydrides), poly(trimethylene carbonate), and two different poly(iminocarbonates). For each polymer, the thermal properties (glass transition temp., crystn., melting and decompn. points) were detd. by DSC and by TGA. The tensile properties (Young's modulus, tensile strength, and elongation at yield and break) were detd. by tensile testing on an Instron stress-strain tester. The flexural storage modulus as a function of temp. was detd. by dynamic mech. anal.

ΙT 133063-34-0

RL: PRP (Properties)

(biodegradable, physicomech. properties of, for medical applications)

133063-34-0 HCAPLUS RN

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, homopolymer CN (9CI) (CA INDEX NAME)

CM

CRN 133063-33-9

CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

IT 133063-34-0

RL: PRP (Properties)

(biodegradable, physicomech. properties of, for medical applications)

L92 ANSWER 41 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1992:557568 HCAPLUS

DN 117:157568

TI Tissue compatibility of tyrosine-derived polycarbonates and polyiminocarbonates: an initial evaluation

AU Silver, Frederick H.; Marks, Michael; Kato, Yasushi P.; Li, Chun; Pulapura, Satish; Kohn, Joachim

CS Biomater. Cent., Robert Wood Johnson Med. Sch., Piscataway, NJ, 08854, USA

SO J. Long-Term Eff. Med. Implants (1992), 1(4), 329-46 CODEN: JLEIEM; ISSN: 1050-6934

DT Journal

LA English

AB

Compression-molded disks of two tyrosine-derived polymers [poly(deaminotyrosyl-tyrosine-hexyl ester carbonate) and poly(deaminotyrosyl-tyrosine hexyl ester iminocarbonate)], 2 polymers derived from Bisphenol A [poly(Bisphenol A iminocarbonate) and poly(Bisphenol A N-phenyliminocarbonate)], and 2 clin. used std. materials [poly(DL-lactic acid) and high-d. polyethylene] were implanted s.c. in back of Sprague-Dawley rats. The tissue response elicited by these materials was evaluated histol. at 7, 30, and 120 days postimplantation, based on the total cell d. (including fibroblasts, monocytes, giant cells, and macrophages) at the implantation site. The tissue response obsd. for the two tyrosine-derived polymers was mild, comparable to the 2 std. materials, medical-grade poly(L-lactic acid) and high d. polyethylene. The 2 Bisphenol A-contq. polymers elicited significantly more severe tissue responses. The use of derivs. of the natural amino acid 1-tyrosine in the synthesis of degradable implant materials improved the tissue compatibility of these materials relative to chem. related polymers that contain Bisphenol A, an industrial diphenol. The tyrosine-derived polyiminocarbonate and polycarbonate are therefore promising candidates for a detailed evaluation of their biocompatibility, including long-term implantation studies in higher mammals.

IT 143715-03-1

RL: MSC (Miscellaneous)

(tissue compatibility of, for prosthetic implants)

RN 143715-03-1 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic acid (9C1) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5

CDES 5:L

Absolute stereochemistry.

CM

CRN 463-79-6 CMF C H2 O3

- C-- OH

IT 143715-03-1 143715-04-2

RL: MSC (Miscellaneous)

(tissue compatibility of, for prosthetic implants)

ANSWER 42 OF 54 HCAPLUS COPYRIGHT 2001 ACS L92

1992:470492 HCAPLUS AN

DN 117:70492

ΤI Amino acid-derived bisphenols for bioerodible polymers

ΙN Kohn, Joachim B.; Pulapura, Satish K. K.

Rutgers, State University, USA PA

U.S., 7 pp. SO

CODEN: USXXAM

DT Patent

LA English

FAN.CNT 2									
PATENT	NO. KIN	D DATI	DATE		LICATION NO.	DATE			
PI US 509	9060 A	1992	20324	US	1990-536425	19900612	<		
US 519	8507 A	1993	30330	US	1991-804767	19911209	<		
US 521	6115 A	1993	30601	US	1992-930146	19920813	<		
US 531	7077 A	199	40531	US	1993-39929	19930329	<		
PRAI US 199	0-536425 199	00612	<						
US 199	1-804767 199	11209	<						
US 199	2-930146 199	20813	<						
OS MARPAT 117:70492									

The title bisphenols HOC6H4CH2CH2CONHCH(CO2R1)CH2C6H4OH (R = C1-18 alkyl, AB esp. hexyl) are useful in the manuf. of polymers (e.g., polycarbonates) which are compatible with living tissue and useful as biomedical prostheses and implants susceptible to gradual erosion within the body.

ΙT 133418-81-2P

RL: PREP (Preparation)

(prepn. of biodegradable, for bioprostheses and implants)

RN 133418-81-2 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C C12 O

## IT 133418-81-2P

RL: PREP (Preparation)

(prepn. of biodegradable, for bioprostheses and implants)

L92 ANSWER 43 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1992:455888 HCAPLUS

DN 117:55888

TI Tyrosine-derived polycarbonates: backbone-modified "pseudo"-poly(amino acids) designed for biomedical applications

AU Pulapura, Satish; Kohn, Joachim

CS Dep. Chem., Rutgers, State Univ., New Brunswick, NJ, 08855-0939, USA

SO Biopolymers (1992), 32(4), 411-17

CODEN: BIPMAA; ISSN: 0006-3525

DT Journal

LA English

AB

Starting from L-tyrosine (Tyr) and its metabolites desaminotyrosine (Dat) and tyramine (Tym), four structurally related model dipeptides were prepd.: Dat-Tym (neither N- or C-terminus present), Z-Tyr-Tym (N-terminus protected by benzyloxycarbonyl) Dat-Tyr-Hex (C-terminus protected by a hexyl ester group), and Z-Tyr-Tyr-Hex (both N- and C-termini present, protected by benzyloxycarbonyl and hexyl ester, resp.). The model dipeptides were used a monomers in the synthesis of polycarbonates. The polymn. reaction in the presence of either phosgene or triphosgene proceeded via the phenolic hydroxyl groups. Polymers with mol. wts. of 105,000-400,000 Datlon (by gel permeation chromatog., relative to polystyrene stds.) were obtained. The physicomech. properties (soly., mech. strength, glass transition and decompn. temp., processibility) of the polymers were detd., and an attempt was made to correlate the polymer properties with the nature of the N- and C-terminus protecting groups. The presence of the urethane bond at the N-terminus protecting group was found to reduce soly., ductility, and processibility, probably due to interchain hydrogen bonding. The presence of a C-terminus alkyl ester group increased soly. and processibility. Thus, the most promising candidate polymer for biomedical applications was obtained from Dat-Tyr-Hex, the monomer carrying a C-terminus protecting group only. Since very similar results had recently been obtained for a series of structurally related polyiminocarbonates, the structure property correlations seems to be generally valid.

IT 133063-34-0P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and physicomech. properties of, structure effect on, biomaterials in relation to)

RN 133063-34-0 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

## IT 133063-34-0P 133063-35-1P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and physicomech. properties of, structure effect on, biomaterials in relation to)

L92 ANSWER 44 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1992:201112 HCAPLUS

DN 116:201112

TI Polyalkylene oxide-amino acid copolymers as drug carriers and charged copolymers based thereon

IN Zalipsky, Samual; Bolikal, Durgadas; Nathan, Aruna; Kohn, Joachim

Benjamin

PA Enzon, Inc., USA

SO PCT Int. Appl., 97 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 2

	PA	TENT 1	NO.		KIN	1D	DATE			APPI	LICATIO	ON NO.	DATE	
ΡI	WO	9200	748		<b>A</b> 1	_	1992	0123		WO 1	L991-U	S4797	199107	708 <
		W:	AU,	CA,	HU,	JP,	SU							
		RW:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB, GF	R, IT,	LU, N	NL, SE	
	JΡ	05508	3879		Т2	2	1993	1209		JP 1	1991-5	12668	199107	708 <
PRAI	US	1990-	-5494	194	199	007	06	<						
	US	1991-	-7263	301	199	107	05	<						
	WO	1991-	-US47	797	199	107	08	<						
		_												

AB Copolymers of polyalkylene oxides and amino acids or peptide sequences are disclosed, which amino acids or peptide sequences have pendant functional groups that are capable of being conjugated with pharmaceutically active compds. for drug delivery systems and crosslinked to form polymer matrixes as hydrogel membranes. The copolymers can also be formed into conductive materials by combination with electrolyte salts. Thus, polyethylene glycol-lysine copolymer was treated with N-hydroxysuccinimide and dicyclohexyl carbodiimide. Cephradine dissolved in a water-dioxane mixt. was reacted with the derivatized polyethylene glycol-lysine copolymer to prep. a conjugate.

IT 133418-81-2

RL: BIOL (Biological study)

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

RN 133418-81-2 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5

CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C Cl2 O

## IT 133418-81-2

RL: BIOL (Biological study)

(semi-interpenetrating polymer networks contg. polyalkylene oxide-amino acid copolymers and, for medical goods)

L92 ANSWER 45 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1991:186204 HCAPLUS

DN 114:186204

TI The synthesis and characterization of pseudopoly(amino acids): new polymers for medical applications

AU Kohn, J.

CS Dep. Chem., Rutgers, State Univ., New Brunswick, NJ, 08855, USA

SO Polym. Prepr. (Am. Chem. Soc., Div. Polym. Chem.) (1990), 31(2),

CODEN: ACPPAY; ISSN: 0032-3934

DT Journal

LA English

N-Hexadecanoyl-hydroxy-L-proline Me ester is prepd. and polymd. by melt polycondensation to yield a polyester with pendent amide groups. The analogous reaction with N-protected-L-serine fails because of .beta.-elimination; the amine group is therefore protected with a benzyloxycarbonyl group, and a poly(serine ester) is obtained by ring-opening polymn. of the serine .beta.-lactone. A diol is prepd. from desaminotyrosine(3-(4-hydroxyphenyl)propanoic acid) and tyrosine hexyl ester, and is then polycondensed with phosgene to yield a polycarbonate-polyamide with pendent ester groups. The biocompatibility and hydrolysis of these polymers in the human body and their use in

medical applications is discussed.

IT 133418-81-2P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. and use of, in medical applications)

RN 133418-81-2 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C C12 O

### IT 133418-81-2P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and use of, in medical applications)

L92 ANSWER 46 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1991:171127 HCAPLUS

DN 114:171127

TI Structure-property relationships for the design of polyiminocarbonates

AU Pulapura, Satish; Li, Chun; Kohn, Joachim

CS Dep. Chem., Rutgers, State Univ., New Brunswick, NJ, USA

SO Biomaterials (1990), 11(9), 666-78

CODEN: BIMADU; ISSN: 0142-9612

DT Journal

LA English

AB Structure-property relationships for the design of new polyiminocarbonates were established, based on the investigation of thermal stability and processibility, morphol., tensile strength, hydrolytic degrdn. and drug release profiles of 15 different polyiminocarbonates. Some polyiminocarbonates were among the mech. strongest, bioerodible polymers currently available. The iminocarbonate bond was highly unstable under physiol. conditions, facilitating the design of rapidly degrading devices. The drug-release profiles of certain polyiminocarbonates exhibited lag periods, facilitating the design of pulsed-release or delayed-release devices. Possible limitations of the practical applicability of polyiminocarbonates as biomaterials were the low thermal stability of the iminocarbonate linkage and the complicated, two-phase degrdn. mechanism

that led to the formation of slowly degrading residues of low mol. wt. To identify non-toxic diphenols as monomers for the synthesis of polyiminocarbonates, derivs. of tyrosine dipeptide were systematically explored. Using structure-property relationships as design guidelines, desaminotyrosyl-tyrosine hexyl ester was identified as a promising, tyrosine-derived diphenol. The corresponding poly(desaminotyrosyl-tyrosine hexyl ester iminocarbonate) formed amorphous, transparent films and was moldable at about 70.degree.. It had a tensile strength of 400 kg/cm2 and a tensile modulus of 16,300 kg/cm2. Under physiol. conditions in vitro, a thin film made of high mol. wt. poly(desaminotyrosyl-tyrosine hexyl ester iminocarbonate) degraded to low mol. wt. oligomers with 5 days. Thus, polyiminocarbonates and in particular poly(desaminotyrosyl-tyrosine hexyl ester iminocarbonate) might be of interest in a variety of biomedical applications.

IT 118798-95-1P

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. and structure-property relationships and drug release from)

RN 118798-95-1 HCAPLUS

CN L-Tyrosine, N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, polymer with (1-methylethylidene)di-4,1-phenylene dicyanate (9CI) (CFINDEX NAME)

CM 1

CRN 106231-84-9 CMF C32 H38 N2 O7 CDES 5:L,L

Absolute stereochemistry.

CM 2

CRN 1156-51-0 CMF C17 H14 N2 O2

IT 118798-95-1P 128835-54-1P 130005-63-9P 133063-34-0P 133063-35-1P 133063-37-3P 133063-38-4P 133134-42-6P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and structure-property relationships and drug release from)

L92 ANSWER 47 OF 54 HCAPLUS COPYRIGHT 2001 ACS AN 1990:578192 HCAPLUS

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DN 113:178192
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TI Biomaterials based on "pseudo"-poly(amino acids): a study of tyrosine-derived polyiminocarbonates

AU Pulapura, S.; Kohn, J.

CS Dep. Chem., Rutgers Univ., New Brunswick, NJ, 08855, USA

SO Polym. Prepr. (Am. Chem. Soc., Div. Polym. Chem.) (1990), 31(1), 233-4 CODEN: ACPPAY; ISSN: 0032-3934

DT Journal

LA English

GΙ

AB Polyiminocarbonates, e.g. I (R = Et, hexyl, or palmityl) were prepd. and their properties detd. Incorporation of nonamide linkages into the backbone of the poly(amino acids) leads to an improvement of the processibility and the physicomech. properties of the polymers. None of the polymers exhibited gross toxicity or tissue incompatibility on s.c. implantation in mice, rats, or rabbits.

IT 106231-87-2P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and properties of, for biomaterials)

RN 106231-87-2 HCAPLUS

CN L-Tyrosine, N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, 4-carbamate, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 106231-86-1 CMF C33 H39 N3 O8 CDES 5:L,L

Absolute stereochemistry.

### IT 106231-87-2P 128835-54-1P 128835-60-9P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and properties of, for biomaterials)

L92 ANSWER 48 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1990:119400 HCAPLUS

DN 112:119400

TI Backbone modification of synthetic poly-.alpha.-L-amino acids

AU Kohn, Joachim; Langer, Robert

CS Dep. Chem., Rutgers, State Univ., Piscataway, NJ, 08855-0939, USA

SO Pept.: Chem. Biol., Proc. Am. Pept. Symp. 10th (1988), Meeting Date 1987,

658-61. Editor(s): Marshall, Garland R. Publisher: ESCOM Sci. Pub.,

Leiden, Neth. CODEN: 56MDA6 Conference

English

LA GI

DT

$$\begin{array}{c|c} & \text{CO}_2R & \text{NH} \\ | & \text{OCO} \\ & \text{NHCO}_2\text{CH}_2\text{Ph} \\ & & \text{NH} \end{array}$$

AB A symposium on the prepn. of tyrosyltyrosine side chain polymers I (R = Et, hexyl, palmityl). The polymers show promise as biodegradable implantable drug delivery devices.

IT 93174-02-8P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn., melting range, and soly. of, in methylene chloride)

RN 93174-02-8 HCAPLUS

CN L-Tyrosine, N-[O-cyano-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, ethyl ester, cyanate (ester), polymer with N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-L-tyrosine ethyl ester (9CI) (CA INDEX NAME)

CM 1

CRN 93174-01-7 CMF C30 H28 N4 O7 CDES 5:L,L

Absolute stereochemistry.

CM 2

CRN 4142-95-4 CMF C28 H30 N2 O7 CDES 5:L,L

# IT 93174-02-8P 118949-20-5P 125607-28-5P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn., melting range, and soly. of, in methylene chloride)

L92 ANSWER 49 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1989:595409 HCAPLUS

DN 111:195409

TI Preparation of nonpeptide polyamino acid bioerodible polymers for drug formulations

IN Kohn, Joachim; Langer, Robert S.

PA Massachusetts Institute of Technology, USA

SO U.S., 9 pp. CODEN: USXXAM

DT Patent

LA English

FAN, CNT 2

11111	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	US 4638045	Α	19870120	US 1985-703153	19850219
	US 4863735	Α	19890905	US 1986-914380	19861002
PRAT	US 1985-703153	19850219			

Biodegradable polymers are prepd. by polymn. of ZNHCHR1CONHCHR2COY or ZNHCHR1CONHCHR2CONHCHR3COY (R1-R3 = side chains of L-.alpha.-amino acids; Y, Z = protecting group) through .gtoreq.1 of R1-R3, useful for controlled release applications in vivo and vitro for delivery of a wide variety of biol. and pharmacol. active ligands. are prepd. Thus, Z-Glu-Phe-OH (Z = PhCH2O2C) and Et3N in CH2Cl2 were treated with (PhO)2P(O)Cl and the mixt. was kept at 4.degree. for 12 h to give a polymer with an av. mol. wt. of 17,000.

IT 123375-14-4P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of, as bioerodible material)

RN 123375-14-4 HCAPLUS

CN L-Tyrosine, N-[O-(aminocarbonyl)-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, ethyl ester, carbamate (ester), homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 123375-13-3 CMF C30 H32 N4 O9 CDES 5:L,L

### IT 123375-14-4P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of, as bioerodible material)

L92 ANSWER 50 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1989:213501 HCAPLUS

DN 110:213501

TI Synthesis of poly(iminocarbonates): degradable polymers with potential applications as disposable plastics and as biomaterials

AU Li, Chun; Kohn, Joachim

CS Dep. Chem., Rutgers, State Univ., New Brunswick, NJ, 08903, USA

SO Macromolecules (1989), 22(5), 2029-36 CODEN: MAMOBX; ISSN: 0024-9297

DT Journal

LA English

The prepn. of poly(iminocarbonates) by bulk, soln., and interfacial AΒ polycondensation was studied by model copolymn. of bisphenol A (I) with bisphenol A dicyanate (II). Bulk polymn. was unsuitable for prepn. of structurally well-defined poly(iminocarbonates) due to the formation of crosslinked products. The soln. polymn. of I with II was optimized in terms of solvent, catalyst, and catalyst concn. In THF with K tert-butoxide as the catalyst, polymers with mol. wts. up to 80,000 were obtained. Interfacial polymn. was a feasible prepn. technique in spite of the high sensitivity of cyanates toward hydrolysis. In the presence of Bu4NBr as a phase-transfer catalyst, polymers with mol. wts. >100,000 were obtained. The interfacial polymn. of I with CNBr instead with II was also feasible, eliminating the need to isolate and purify the reactive dicyanate. Novel poly(iminocarbonates) were prepd. by soln. polymn. of II with 4,4'-thiodiphenol, desaminotyrosyl tryramine, or N-(benzoyloxy) carbonyl-L-tyrosyl-L-tyrosine hexyl ether. The potential applications of poly(iminocarbonates) as disposable plastics and biomaterials were discussed.

IT 118798-95-1P, N-(Benzyloxy)carbonyl-L-tyrosyl-L-tyrosine hexyl
 ester-bisphenol A dicyanate copolymer
 RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. and characterization of)

RN 118798-95-1 HCAPLUS

CN L-Tyrosine, N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester,
 polymer with (1-methylethylidene)di-4,1-phenylene dicyanate (9CI) (CA
 INDEX NAME)

CM 1

CRN 106231-84-9 CMF C32 H38 N2 O7 CDES 5:L,L

CM 2

CRN 1156-51-0 CMF C17 H14 N2 O2

L92 ANSWER 51 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1989:101818 HCAPLUS

DN 110:101818

TI Implantable biodegradable polymeric drug delivery system with adjuvant activity

IN Kohn, Joachim B.; Langer, Robert S.; Niemi, Steven M.; Fox, James G.

PA Massachusetts Institute of Technology, USA

SO PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN. CNT 2

PAN.	CNT Z			
	PATENT NO.	KIND DATE	APPLICATION NO.	DATE
ΡT	WO 8802262	A1 19880407	WO 1987-US2428	19870924
	W: JP	111 13000107	WO 1307 ODE 120	150.0511
	RW: AT, BE,	CH, DE, FR, GB, IT	C, LU, NL, SE	
	US 4863735	A 19890905	US 1986-914380	19861002
	CA 1323566	A1 19931026	CA 1987-548151	19870929
PRAI	US 1986-914380	19861002	•	
	US 1985-703153	19850219	,	

AB An antigen delivery system is described which utilizes a biodegradable polymer with good mech. properties in combination with a material stimulating the immune system. The material having adjuvant activity maybe a polymer degrdn. product or an adjuvant which is contained within or bound to the polymer. The polymer may be formed from tyrosine dipeptides. This polymer is not an adjuvant, but degrades into products which stimulate the immune system. The advantages of this system are that a polymer can be used to form a biodegradable integral structure which is useful as both an implantable source of an antigen or other biol. active compds. and as a control means for the rate of release of the biol.

compd., producing a sustained, relatively const. delivery of antigen with simultaneous stimulation of the immune response. N-Cbz-Tyr-Tyr-Hex (CTTH; Cbz = benzyloxycarbonyl; Hex = hexyl) (prepn. given) was treated with CNBr to form the dicyanate, which was mixed with an equimolar quantity of CTTH and the mixt. polymd. in the presence of KOBu-tert to give poly(CTTH-iminocarbonate) with mol. wt. 19,500 and which had an intrinsic viscosity 0.27 (DMF, 25.degree.). Using particulate suspensions, the degrdn. products of poly(CTTH-iminocarbonate) were shown to be as potent an adjuvant as complete Freund's adjuvant and muramyl dipeptide when the serum antibody response to bovine serum albumin (BSA) in male CD-1 mice is measured over 56 wk. Further, BSA released from s.c. implanted polymeric antigen delivery devices made of poly(CTTH-iminocarbonate) resulted in significantly higher anti-BSA titers than devices made of poly(bisphenol A-iminicarbonate). At the end of week 56 the injection and implantation sites were examd. and no tissue abnormalities were found, no residues of the injected particulate material were detected, and only small amts. of polymeric residue were detected at the implantation sites.

IT 118949-20-5P

RL: PREP (Preparation)

(manuf. of, as biodegradable polymer for stimulation of immune response)

RN 118949-20-5 HCAPLUS

CN L-Tyrosine, N-[O-cyano-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, cyanate (ester), polymer with N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-L-tyrosine hexyl ester (9CI) (CA INDEX NAME)

CM 1

CRN 106231-84-9 CMF C32 H38 N2 O7 CDES 5:L,L

Absolute stereochemistry.

CM 2

CRN 104549-44-2 CMF C34 H36 N4 O7 CDES 5:L,L

### IT 118949-20-5P

RL: PREP (Preparation)

(manuf. of, as biodegradable polymer for stimulation of immune response)

L92 ANSWER 52 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1987:162477 HCAPLUS

DN 106:162477

TI Single-step immunization using a controlled release, biodegradable polymer with sustained adjuvant activity .

AU Kohn, Joachim; Niemi, Steven M.; Albert, Elizabeth C.; Murphy,

James C.; Langer, Robert; Fox, James G.

CS Dep. Appl. Biol. Sci., Massachusetts Inst. Technol., Cambridge, MA, 02139, USA

SO J. Immunol. Methods (1986), 95(1), 31-8

CODEN: JIMMBG; ISSN: 0022-1759

DT Journal

LA English

GΙ

The use of a biodegradable polymer for antigen delivery based on poly(CTTH-iminocarbonate) (I) [106755-33-3] was investigated. This polymer was selected since its primary degrdn. product, N-benzyloxycarbonyl-L-tyrosyl-L-tyrosine hexyl ester (CTTH) [106231-84-9] was as potent an adjuvant as complete Freund's adjuvant and muramyl dipeptide, when measuring the serum antibody response to bovine serum albumin (BSA) in male CD-1 mice over 56 wk. BSA released from s.c. implanted polymeric antigen delivery devices made of I resulted in significantly higher anti-BSA antibody titers than devices made of poly(bisphenol A-iminocarbonate) [26101-32-6].

IT 107173-10-4

RL: BIOL (Biological study)

(biodegradable, with sustained adjuvant activity)

RN 107173-10-4 HCAPLUS

CN L-Tyrosine, N-[O-cyano-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, cyanate (ester), homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 104549-44-2

CMF C34 H36 N4 O7 CDES 5:L,L

### Absolute stereochemistry.

### IT 107173-10-4

RL: BIOL (Biological study)

(biodegradable, with sustained adjuvant activity)

L92 ANSWER 53 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1987:67663 HCAPLUS

DN 106:67663

TI Polymerization reactions involving the side chains of .alpha.-L-amino acids

AU Kohn, Joachim; Langer, Robert

CS Dep. Appl. Biol. Sci., Massachusetts Inst. Technol., Cambridge, MA, 02139, USA

SO J. Am. Chem. Soc. (1987), 109(3), 817-20

CODEN: JACSAT; ISSN: 0002-7863

DT Journal

LA English

OS CASREACT 106:67663

GI

AB Hydroxyproline deriv. I (Pal = palmitoyl) underwent melt transesterification in the presence of Al isopropoxide to give side-chain polymer II. Z-Tyr-Tyr-OHex (III; Z = PhCH2O2C, Hex = hexyl) was treated with cyanogen bromide to give dicyanate IV. The soln. polymn. of equimolar amts. of III and IV in THF contg. KOCMe3 gave the corresponding iminocarbonate side-chain polymer.

IT 106231-87-2P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of)

RN 106231-87-2 HCAPLUS

CN L-Tyrosine, N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, 4-carbamate, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 106231-86-1 CMF C33 H39 N3 O8 CDES 5:L,L

Absolute stereochemistry.

IT 106231-87-2P

L92 ANSWER 54 OF 54 HCAPLUS COPYRIGHT 2001 ACS

AN 1984:616333 HCAPLUS

DN 101:216333

TI A new approach to the development of bioerodible polymers for controlled release applications employing naturally occurring amino acids

AU Kohn, Joachim; Langer, Robert

CS Whitaker Coll. Health Sci., Massachusetts Inst. Technol., Cambridge, MA, 02139, USA

SO Polym. Mater. Sci. Eng. (1984), 51, 119-21

CODEN: PMSEDG

DT Journal

LA English

AB Dipeptides, e.g., N-carbobenzoxytyrosyltyrosine Et ester (I) [4142-95-4], can be used as monomers for the formation of nonpeptide polymers for controlled-release of drugs. I was prepd. by known methods having 2 reactive, arom. OH groups which could be used for formation of a hydrolytically labile iminocarbonate linkage. I-di-O-cyano-I copolymer [93174-02-8] erodes completely within 93 days when exposed to 0.1M phosphate buffer (pH 7.4) at 37.degree..

IT 93174-02-8P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and biodegradability of, controlled-release applications in relation to)

RN 93174-02-8 HCAPLUS

CN L-Tyrosine, N-[O-cyano-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, ethyl ester, cyanate (ester), polymer with N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-L-tyrosine ethyl ester (9CI) (CA INDEX NAME)

CM 1

CRN 93174-01-7 CMF C30 H28 N4 O7 CDES 5:L,L

CM 2

CRN 4142-95-4 CMF C28 H30 N2 O7 CDES 5:L,L

Absolute stereochemistry.

### IT 93174-02-8P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and biodegradability of, controlled-release applications in relation to)

=> d 193 bib abs fhitstr hitrn tot

L93 ANSWER 1 OF 29 HCAPLUS COPYRIGHT 2001 ACS

AN 2000:676553 HCAPLUS

DN 134:105786

TI Hydrolytic degradation of tyrosine-derived polycarbonates, a class of new biomaterials. Part II: 3-yr study of polymeric devices

AU Tangpasuthadol, V.; Pendharkar, S. M.; Peterson, R. C.; Kohn, J.

CS Department of Chemistry, Rutgers-The State University of New Jersey, Piscataway, NJ, 08854, USA

SO Biomaterials (2000), 21(23), 2379-2387 CODEN: BIMADU; ISSN: 0142-9612

PB Elsevier Science Ltd.

DT Journal

LA English

The kinetics and mechanisms of in vitro degrdn. of tyrosine
-derived polycarbonates, a new class of polymeric
biomaterials, were studied extensively at 37.degree.C. These
polymers carry an alkyl ester pendent chain that allows the
fine-tuning of the polymer's material properties, its biol.
interactions with cells and tissue, and its degrdn. behavior. The
polymer carrying an Et ester pendent chain, poly(DTE carbonate),
has been established as a promising orthopedic implant material,
exhibiting bone apposition when in contact with hard tissue.

Tyrosine-derived polycarbonates are relatively stable and degrade only very slowly in vitro. Therefore, accelerated studies were conducted at 50 and 65.degree.C to observe the behavior of polymers during the later stages of degrdn. Varying the pendent chain length affected the rate of water uptake, initial degrdn. rate, and phys. stability of the polymeric devices. During the 3-yr study, the polymer degraded by random chain cleavage of the carbonate bonds, accompanied by a relatively small amt. of pendent chain de-esterification. No mass loss was obsd. during this period at 37.degree.C, but mass loss was readily evident during the accelerated studies at 50 and 65.degree.C. Thus, it is reasonable to assume that mass loss will occur also at 37.degree.C, albeit only after extensive backbone carbonate cleavage and pendent chain ester hydrolysis. The dimension and surface area of the devices influenced the initial degrdn. rate, but did not significantly affect the overall rate of degrdn. No evidence of "acid dumping" or the release of acidic residues found during the degrdn. of poly(d, 1-lactic acid) were obsd. for this family of tyrosine -derived polycarbonates.

IT 183480-53-7P

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(hydrolytic degrdn. of tyrosine-derived polycarbonates as a class of new biomaterials)

RN 183480-53-7 HCAPLUS

Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-(butoxycarbonyl)-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

### IT 183480-53-7P 183480-54-8P 319916-60-4P

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(hydrolytic degrdn. of tyrosine-derived polycarbonates as a class of new biomaterials)

RE.CNT 16

RE

CN

- (2) Choueka, J; J Biomed Mater Res 1996, V31, P35 HCAPLUS
- (3) Ertel, S; J Biomed Mater Res 1994, V28, P919 HCAPLUS
- (4) Ertel, S; J Biomed Mater Res 1995, V29(11), P1337 HCAPLUS
- (5) Ghorbel, I; J Appl Polym Sci 1995, V55, P173 HCAPLUS
- (6) Grizzi, I; Biomaterials 1995, V16(4), P305 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L93 ANSWER 2 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 2000:676552 HCAPLUS
- DN 134:105785
- TI Hydrolytic degradation of tyrosine-derived polycarbonates, a class of new biomaterials. Part I: Study of model compounds
- AU Tangpasuthadol, V.; Pendharkar, S. M.; Kohn, J.
- CS Department of Chemistry, Rutgers-The State University of New Jersey, Piscataway, NJ, 08854-8087, USA
- SO Biomaterials (2000), 21(23), 2371-2378 CODEN: BIMADU; ISSN: 0142-9612

PB Elsevier Science Ltd.

DT Journal

LA English

Tyrosine-derived polycarbonates have been identified AB as promising, degradable polymers for use in orthopedic applications. These polymers are non-toxic, biocompatible, and exhibit good bone apposition when in contact with hard tissue. Tyrosine-derived polycarbonates were designed to incorporate two hydrolytically labile bonds in each repeat unit, a carbonate bond that connects the monomer units and an ester bond connecting a pendent chain. The relative hydrolysis rate of the two bonds will det. the type of degrdn. products and the degrdn. pathway of the polymers. In order to study the degrdn. mechanism of these polycarbonates in more detail, a series of small model compds. were designed that mimic the repeat unit of the polymer. Results obtained from the use of these model compds. suggested that the backbone carbonate bond is hydrolyzed at a faster rate than the pendent chain ester bond. Increasing the length of the alkyl pendent chain lowered the hydrolysis rates of both hydrolyzable linkages, possibly by hindering the access of water mols. to those sites. The hydrolysis rates of both linkages were pH dependent with the lowest rate at pH about 5. The results from this study can be used to explain the degrdn. behavior of the corresponding polycarbonates as well as their degrdn. mechanisms. This information is essential when evaluating the utility of tyrosine-derived polycarbonates as degradable medical implant materials.

IT 183480-55-9P

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(hydrolytic degrdn. of tyrosine-derived polycarbonate as a class of biomaterials)

RN 183480-55-9 HCAPLUS

Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-[(octyloxy)carbonyl]-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

IT 183480-55-9P

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(hydrolytic degrdn. of tyrosine-derived polycarbonate as a class of biomaterials)

RE.CNT 20

RE

CN

- (1) Chasin, M; Biodegradable polymers as drug delivery systems 1990, P43 HCAPLUS
- (2) Choueka, J; J Biomed Mater Res 1996, V31, P35 HCAPLUS
- (4) Doddi, N; US 4052988 1977 HCAPLUS
- (5) Ertel, S; J Biomed Mater Res 1994, V28, P919 HCAPLUS
- (6) Grizzi, I; Biomaterials 1995, V16(4), P305 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L93 ANSWER 3 OF 29 HCAPLUS COPYRIGHT 2001 ACS

```
ΑN
     2000:568454 HCAPLUS
     133:168440
DN
ΤI
     Porous polymer scaffolds for tissue engineering
     Levene, Howard B.; Lhommeau, Christelle M.; Kohn, Joachim B.
ΙN
·PA
     Rutgers, the State University, USA
SO
     U.S., 11 pp.
     CODEN: USXXAM
DT
     Patent
LΑ
     English
FAN.CNT 1
                        KIND DATE
     PATENT NO.
                                              APPLICATION NO. DATE
                        ----
                              _____
                                              ______
     US 6103255
ΡI
                              20000815
                                              US 1999-293118
                        Α
                                                                 19990416
     WO 2000062829
                              20001026
                       A1
                                              WO 1999-US8375
                                                               19990416
             AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,
              DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS,
              JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,
              MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
              MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
              ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
                      GA, GN, GW, ML, MR, NE, SN, TD, TG
     AU 9935663
                        A1
                             20001102
                                              AU 1999-35663
                                                                 19990416
PRAI US 1999-293118
                        19990416
     WO 1999-US8375
                        19990416
     Biodegradable and biocompatible porous scaffolds characterized by a
AΒ
     substantially continuous polymer phase, having a highly interconnected
     bimodal distribution of open pore sizes with rounded large pores of about
     50 to about 500 .mu.m in diam. and rounded small pores less than 20 .mu.m \,
     in diam., wherein the small pores are aligned in an orderly linear fashion
     within the walls of the large pores. Methods of prepg. polymeric tissue scaffolds are also disclosed. Thus, 0.3 g of poly(L-lactide) was dissolved in 1,4-dioxane/water (91/9% vol./vol.)., the clear soln. was
     then poured on 7 g of sieved sodium chloride salts in a dish. After the
     diffusion of the polymer soln. through the salt bed, it was freeze-dried
     leaving a porous structure. The polymer did not relax during solvent
     removal. Finally, the salt was leached out in water.
                                                                The water was
     changed several times until the sensitive silver nitrate test did not show
     any addnl. release of chloride ions into the water. The resulting
     scaffolds were removed from the water and dried for several days to const.
          The dried scaffolds were very soft and could be easily deformed
     because of the high total porosity and the low polymer modulus.
TΤ
     188712-99-4P
     RL: DEV (Device component use); PNU (Preparation, unclassified); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
         (porous polymer scaffolds for tissue engineering)
RN
     188712-99-4 HCAPLUS
CN
     L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, butyl ester, polymer
     with butanedioic acid (9CI) (CA INDEX NAME)
     CM
          1
     CRN 174702-84-2
     CMF C22 H27 N O5
     CDES 5:L
```

CM 2

CRN 110-15-6 CMF C4 H6 O4

 $HO_2C-CH_2-CH_2-CO_2H$ 

23

# IT 188712-99-4P 191858-68-1P 214259-59-3P 219622-84-1P

RL: DEV (Device component use); PNU (Preparation, unclassified); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(porous polymer scaffolds for tissue engineering)

RE.CNT

- (1) Anon; WO 9425079 1994 HCAPLUS
- (3) Cohn; US 4826945 1989 HCAPLUS
- (4) Degroot; Colloid Polym Sci 1990, V268, P1073 HCAPLUS
- (5) Healy; US 5723508 1998 HCAPLUS
- (6) Kohn; US 5099060 1992 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L93 ANSWER 4 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 2000:226537 HCAPLUS
- DN 133:109861
- TI Characterization of the inflammatory response to biomaterials using a rodent air pouch model
- AU Hooper, Kimberly A.; Nickolas, Thomas L.; Yurkow, Edward J.; Kohn, Joachim; Laskin, Debra L.
- CS Department of Pharmacology and Toxicology, Rutgers, The State University of New Jersey, Piscataway, NJ, 08854-8020, USA
- SO J. Biomed. Mater. Res. (2000), 50(3), 365-374 CODEN: JBMRBG; ISSN: 0021-9304
- PB John Wiley & Sons, Inc.
- DT Journal
- LA English
- Using a rodent air pouch, the inflammatory responses to biomaterials with distinct phys. properties and chem. compns. were compared. The polymers examd. were expanded poly(tetrafluoroethylene) (ePTFE), silicone, low-d. polyethylene (LDPE), poly(L-lactic acid) (PLLA), poly(desaminotyrosyl-tyrosine Et carbonate) [poly(DTE carbonate)], and poly(desaminotyrosyl-tyrosine benzyl carbonate) [poly(DTBzl carbonate)]. We found that implantation of disks (4.5-4.8 mm) of these materials into rodent air pouches for 2 days had no effect on the no. or type of cells recovered relative to sham controls. With each of the materials, macrophages were the predominant cell type identified (60-75%), followed by granulocytes (20-25%) and lymphocytes (10%). Implantation of poly(DTE carbonate),

ePTFE, LDPE, or poly(DTBzl carbonate) into the pouches for 2 days caused an increase in release of superoxide anion by the pouch cells. Cells from pouches contg. poly(DTE carbonate) also released more hydrogen peroxide and were more phagocytic. In contrast, PLLA and silicone had no effect on the functional activity of cells recovered from the pouches. Prolonging the implantation time of poly(DTE carbonate) or PLLA to 7 days did not alter the no. or type of cells isolated from the pouches. However, cells from pouches contg. poly(DTE carbonate) for 7 days continued to produce increased quantities of superoxide anion relative to sham control pouch cells. These results suggest that the air pouch model is a highly sensitive method and therefore useful for evaluating the functional responses of inflammatory cells to biomaterials.

IT 219622-84-1

> RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(characterization of inflammatory response to biomaterials using rodent air pouch model)

RN 219622-84-1 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, ethyl ester, polymer with carbonic dichloride (9CI) (CA INDEX NAME)

CM

CRN 135313-59-6 CMF C20 H23 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 75-44-5 CMF C C12 O

#### TΤ 219622-84-1 219622-85-2

RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(characterization of inflammatory response to biomaterials using rodent air pouch model)

RE.CNT 41

- (2) Anderson, J; Biomaterials 1984, V5, P5 HCAPLUS
- (3) Athanasiou, K; Biomaterials 1996, V17, P93 HCAPLUS
- (6) Behling, C; J Biomed Mater Res 1986, V20, P653 HCAPLUS(7) Bergsma, J; J Biomed Mater Res 1995, V29, P173 HCAPLUS
- (9) Cerami, A; Clin Immunol Immunopath 1992, V62, PS3 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L93 ANSWER 5 OF 29 HCAPLUS COPYRIGHT 2001 ACS

```
2000:225757 HCAPLUS
AN
DN
     132:348250
     Characterization of combinatorially designed polyarylates by
ΤI
     time-of-flight secondary ion mass spectrometry
     Belu, Anna M.; Brocchini, Stephen; Kohn, Joachim;
ΑU
     Ratner, Buddy D.
     Department of Bioengineering, University of Washington, Seattle, WA,
CS
     98195-1750, USA
     Rapid Commun. Mass Spectrom. (2000), 14(7), 564-571
SO
     CODEN: RCMSEF; ISSN: 0951-4198
PΒ
     John Wiley & Sons Ltd.
DT
     Journal
LA
     English
     A series of 16 polyarylates, with well-controlled and systematically
AB
     varying chem., has been characterized by time-of-flight secondary ion mass
     spectrometry (TOF-SIMS). The polymers are structurally
     identical except for the incremental addns. of C2H4 units to the backbone
     and side-chain. From the spectra, peaks characteristic of all
     polyarylates are identified. Furthermore, evaluation of the spectra and
     identification of unique signals allow classification of the polyarylates
     according to side-chain and backbone chem.
ΙT
     149787-38-2
     RL: PRP (Properties)
        (time-of-flight secondary ion mass spectrometry of tyrosine
        ester-based polyesters)
RN
     149787-38-2 HCAPLUS
     L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer
CN
     with butanedioic acid (9CI) (CA INDEX NAME)
```

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

Me (CH<sub>2</sub>) 
$$5$$
 O O OH

CM 2

CRN 110-15-6 CMF C4 H6 O4

 ${\tt HO_2C-CH_2-CH_2-CO_2H}$ 

IT 149787-38-2 149787-39-3 149787-40-6 149787-41-7 149826-02-8 188712-92-7 188712-95-0 188712-97-2 188712-99-4 188713-01-1 188713-03-3 188713-05-5 188713-08-8 188713-09-9 188713-10-2 188713-11-3 RL: PRP (Properties) (time-of-flight secondary ion mass spectrometry of tyrosine
ester-based polyesters)

RE.CNT 5

RE

- (1) Brocchini, S; J Am Chem Soc 1997, V119, P4553 HCAPLUS
- (2) Fiordeliso, J; J Biomater Sci Polym Ed 1994, V5, P497 HCAPLUS
- (3) Perez-Luna, V; J App Polym Sci 1997, V63, P1467 HCAPLUS
- (4) Reichlmaier, S; Surf Interface Anal 1994, V21, P739 HCAPLUS
- (5) Tamada, Y; J Biomed Mater Res 1994, V28, P283
- L93 ANSWER 6 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 2000:108769 HCAPLUS
- DN 132:270023
- TI PEG-variant biomaterials as selectively adhesive protein templates: model surfaces for controlled cell adhesion and migration
- AU Tziampazis, Evangelos; Kohn, Joachim; Moghe, Prabhas V.
- CS Department of Chemical and Biochemical Engineering, Rutgers University, Piscataway, NJ, 08854-8058, USA
- SO Biomaterials (2000), 21(5), 511-520 CODEN: BIMADU; ISSN: 0142-9612
- PB Elsevier Science Ltd.
- DT Journal
- LA English
- AΒ Our study focused on the role of poly(ethylene glycol) (PEG) in actively regulating the biol. responsiveness of protein-adsorbed biomaterials. To this end, we designed PEG-variant biomaterials from a family of tyrosine/PEG-derived polycarbonates to present surfaces ranging from low to intermediate levels of PEG concn., below the PEG level requisite for complete abolition of protein adsorption. We analyzed the effect of PEG concn. on the amt., conformation and bioactivity of an adsorbed model protein, fibronectin, and on the attachment, adhesion strength and motility of L929 fibroblasts. Our results demonstrate that low levels of PEG can regulate not only the extent but also the conformation and specific bioactivity of adsorbed fibronectin. As the PEG concn. was increased from 0 to 6 mol%, the amt. of adsorbed fibronectin decreased linearly yet the fibronectin conformation was altered such that the overall bioactivity of adsorbed fibronectin was uncompromised. We report that the degree of cell attachment varied with PEG concn. in a manner similar to the dependence of fibronectin bioactivity on PEG. In contrast, the nature of cell adhesion strength dependence on PEG paralleled the pattern obsd. for fibronectin surface concn. Our studies also indicated that the rate of cell migration was inversely correlated with PEG concn. over a narrow range of PEG concn. Overall, these results highlight the striking ability of PEG-variant biomaterials to systematically regulate the behavior of adsorbed cell adhesion proteins and, consequently, effect cell functions.

### IT 263565-88-4

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(PEG-variant biomaterials as selectively adhesive protein templates as model surfaces for controlled cell adhesion and migration)

RN 263565-88-4 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, ethyl ester, polymer with .alpha.-carboxy-.omega.-(carboxyoxy)poly(oxy-1,2-ethanediyl) (9CI) (CA INDEX NAME)

CM 1

CRN 135313-59-6 CMF C20 H23 N O5 CDES 5:L

CM 2

CRN 85022-96-4

CMF (C2 H4 O)n C2 H2 O5

CCI PMS

$$HO_2C$$
  $O-CH_2-CH_2$   $O-CO_2H$ 

### IT 263565-88-4

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(PEG-variant biomaterials as selectively adhesive protein templates as

model surfaces for controlled cell adhesion and migration)

RE.CNT 49

RE

- (1) Absolom, D; Biochim Biophys Acta 1981, V670, P74 HCAPLUS
- (2) Absolom, D; J Biomed Mater Res 1987, V21, P161 HCAPLUS
- (3) Amiji, M; Biomaterials 1992, V13, P682 HCAPLUS
- (4) Andrade, J; Interfacial phenomena and bioproducts 1996, P19 HCAPLUS
- (5) Arakawa, T; Biochemistry 1985, V24, P6756 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L93 ANSWER 7 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 2000:5405 HCAPLUS
- DN 132:212631
- TI Small changes in **polymer** chemistry have a large effect on the bone-implant interface: evaluation of a series of degradable **tyrosine**-derived **polycarbonates** in bone defects
- AU James, Kenneth; Levene, Howard; Parsons, J. Russell; Kohn, Joachim
- CS Department of Chemistry, Rutgers, The State University of New Jersey, New Brunswick, NJ, 09803, USA
- SO Biomaterials (1999), 20(23/24), 2203-2212 CODEN: BIMADU; ISSN: 0142-9612
- PB Elsevier Science Ltd.
- DT Journal
- LA English

AB

In a series of homologous, tyrosine-based polycarbonates, small changes in the chem. structure of the polymer pendent chain affectes the bone response in a long-term (1280 days) implantation study. Identically sized pins, prepd. from poly(DTE carbonate), poly(DTB carbonate), poly(DTH carbonate), and poly(DTO carbonate) were implanted transcortically in the proximal tibia and the distal femur of skeletally mature New Zealand White Rabbits. The tissue response at the bone-implant interface was characterized in terms of the absence of a fibrous capsule (direct bone apposition, indicative of a bone bonding response) or the presence of a fibrous capsule (referred to as the encapsulation response). The relative frequency of direct bone apposition vs. encapsulation was recorded for each polymer throughout the entire period of the study. While all 4 polymers were tissue compatible, there was a

correlation between the chem. structure of the pendent chain and the type of bone response obsd., with poly(DTE carbonate) having the highest tendency to elicit direct bone apposition. Based on in vivo degrdn. data and the ability of model polymers with carboxylate groups at their surface to chelate calcium ions, it is proposed that the ability of poly(DTE carbonate) to bond to bone is caused by the facile hydrolysis of the pendent Et ester groups which creates calcium ion chelation sites on the polymer surface. The incorporation of calcium chelation sites into the chem. structure of an implant material appears to be a key requirement if direct bone apposition/bone bonding is desired. Very subtle changes in the chem. compn. of an implant material can have significant effects on the long-term tissue response in a clin. relevant model.

IT 183480-53-7

RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(degradable tyrosine-derived polycarbonates
evaluation in bone defects)

RN 183480-53-7 HCAPLUS

CN Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-(butoxycarbonyl)-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

IT 183480-53-7 183480-54-8 183480-55-9

RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(degradable tyrosine-derived polycarbonates evaluation in bone defects)

RE.CNT 24

- (2) Boyan, B; Biomaterials 1996, V17, P137 HCAPLUS
- (3) Brocchini, S; J Amer Chem Soc 1997, V119(19), P4553 HCAPLUS
- (4) Brocchini, S; J Biomed Mater Res 1998, V42, P66 HCAPLUS
- (5) Choueka, J; J Biomed Mater Res 1996, V31, P35 HCAPLUS
- (7) Ertel, S; J Biomed Mater Res 1994, V28, P919 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L93 ANSWER 8 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 1999:784854 HCAPLUS
- DN 132:93927
- TI Conductivity and high-temperature relaxation of tyrosine-derived polyarylates measured with thermal stimulated currents
- AU Puma, M.; Suarez, N.; Kohn, J.
- CS ENINSEL, Carr. Nac. Hoyo de la Puerta, Instituto de Ingenieria, Sartenejas-Caracas, Venez.
- SO J. Polym. Sci., Part B: Polym. Phys. (1999), 37(24), 3504-3511 CODEN: JPBPEM; ISSN: 0887-6266
- PB John Wiley & Sons, Inc.
- DT Journal
- LA English
- AB Thermal-stimulated polarization and depolarization expts. without blocking electrodes are performed on tyrosine-derived polyarylates with different backbone lengths. The expts. on the different samples are carried out using the same thermal history throughout the entire characterization process. The high-temp. current rise caused by the cond. of the samples

is studied with a simple model that utilizes an approxn. of the Williams-Landel-Ferry (WLF) relaxation time. The cond. data is well reproduced except for temps. well below the glass-transition temp. and for small currents. The glass-transition peak is modeled with a phenomenol. expression valid near Tg, which is able to describe the glass relaxation with a min. no. of parameters. The conduction and the glass-transition relaxation are studied vs. the structural changes for the different samples. It is found that the cond. and the glass-transition temp. shift to lower temps. as the methylene groups in the backbone increase. Furthermore, if the exptl. data is presented as a function of the reduced temp., the shape of the glass-transition relaxation for the different samples is independent of the polymer backbone length.

IT 149787-38-2

RL: PRP (Properties)

(cond. and high-temp. relaxation of tyrosine-derived polyarylates measured with thermal stimulated currents)

RN 149787-38-2 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

 $HO_2C-CH_2-CH_2-CO_2H$ 

### IT 149787-38-2 149787-39-3 149787-40-6 188713-08-8

RL: PRP (Properties)

(cond. and high-temp. relaxation of tyrosine-derived polyarylates measured with thermal stimulated currents)

RE.CNT 17

- (3) Brocchini, S; J Biomed Mater Res 1998, V42, P66 HCAPLUS
- (4) Canadas, J; J Polymer 1998, V39, P2795 HCAPLUS
- (5) Chee, K; J Appl Polym Sci 1987, V33, P1067 HCAPLUS
- (7) Fiordeliso, J; J Biomater Sci Polym Ed 1994, V5, P497 HCAPLUS
- (9) Kohn, J; US 5216115 1993 HCAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L93 ANSWER 9 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 1999:722402 HCAPLUS

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DN 132:313446
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TI Polymer ordering enhances delivery of antithrombotic peptide

AU Schachter, D.; Kohn, J.

CS Department of Chemistry, Rutgers University, Piscataway, NJ, 08855-0939, USA

SO Proc. Int. Symp. Controlled Release Bioact. Mater. (1999), 26th, 625-626 CODEN: PCRMEY; ISSN: 1022-0178

PB Controlled Release Society, Inc.

DT Journal

LA English

AB Strong peptide-polymer interactions can occur when formulating a peptide into tyrosine-derived polyarylates because of the peptide-like structure. In the case of polymers that end to order, these interactions remaine strong only in the amorphous regions. In the cryst. regions, the self assocn. of the polymer chains excludes these interactions allowing diffusion to occur.

IT 149787-39-3

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(polymer ordering enhances delivery of antithrombotic peptide)

RN 149787-39-3 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with hexanedioic acid (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

Me 
$$(CH_2)_5$$
 O O OH  $N$  S

CM 2

CRN 124-04-9 CMF C6 H10 O4

 $HO_2C^-$  (CH<sub>2</sub>)<sub>4</sub> - CO<sub>2</sub>H

# IT 149787-39-3 149826-02-8 188712-97-2 188713-11-3

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(polymer ordering enhances delivery of antithrombotic peptide)

RE.CNT 5

- (1) Brocchini, S; JACS 1997, V119, P19
- (2) Gombotz, W; Bioconjugate Chem 1995, V6 HCAPLUS
- (3) Schachter, D; Proc of CRS Conf on Adv in Control 1996
- (4) Schachter, D; manuscript in preparation
- (5) Tcheng, J; Circulation 1995, V91, P8

```
ANSWER 10 OF 29 HCAPLUS COPYRIGHT 2001 ACS
L93
     1999:146214 HCAPLUS
ΑN
ΤТ
     The use of combinatorial approaches in the design of new biomaterials
ΑU
     Kohn, J.; Brocchini, S.; James, K.;
     Tangpasuthadol, V.
CS
     Department of Chemistry, Rutgers University, Piscataway, NJ, 08854, USA
SO
     Book of Abstracts, 217th ACS National Meeting, Anaheim, Calif., March
     21-25 (1999), POLY-178 Publisher: American Chemical Society, Washington,
     CODEN: 67GHA6
DT
     Conference; Meeting Abstract
LA
     English
AΒ
     The development of combinatorial approaches to polymer design provides new
     strategies for the rapid identification of novel biomaterials. Our prior
     experience with the design of polymeric biomaterials indicated that the
     traditional methods of synthesizing new polymers one after the other in a
     sequential fashion are too time consuming to satisfy the materials need of
     emerging fields such as "tissue engineering". To address this problem, we
     have created the first combinatorial library of biomaterials. In this
     library of 112 structurally related polymers, it has been possible to
     identify new correlations between polymer structure, polymer properties,
     and the cellular response in vitro. Data will be presented that
     illustrate the value of this combinatorial approach in identifying useful
     new polymeric compns. for a range of medical applications.
L93
     ANSWER 11 OF 29 HCAPLUS COPYRIGHT 2001 ACS
     1999:109959 HCAPLUS
ΑN
     130:297273
DN
TI
     Tyrosine-PEG-derived poly(ether carbonate)s as new biomaterials. Part II:
     study of inverse temperature transitions
ΑU
     Yu, Chun; Mielewczyk, Slawomir S.; Breslauer, Kenneth J.; Kohn,
     Joachim
CS
     Department of Chemistry, Rutgers, The State University of New Jersey, New
     Brunswick, NJ, 08903, USA
SO
     Biomaterials (1999), 20(3), 265-272
     CODEN: BIMADU; ISSN: 0142-9612
PΒ
     Elsevier Science Ltd.
DT
     Journal
     English
LA
AΒ
     Tyrosine-poly(alkylene oxide)-derived poly(ether carbonates) represent a
     new group of degradable biomaterials that exhibit inverse temp.
     transitions.
                  Poly(DTE co 70%PEG1000 carbonate) (DTE =
     deaminotyrosyltyrosine Et ester) was chosen as an example to study this
     special phase transition behavior of the polymers. The obsd. transition
     temp. varied slightly depending on the technique used, e.g. CD spectra
     always gave a lower temp. than UV/visible spectra. The CD and UV/visible
     studies indicated that the transition temp. was both heating rate and
     concn. dependent. Thermodn. parameters of the transition (enthalpy,
     entropy, and free energy) were detd. by DSC. The molecularity of the
     transition was 2.6, as calcd. from UV and DSC data. The transition temp.
     could be varied from 18 to 580C by changing the polymer structure.
IT
     223114-10-1
     RL: PRP (Properties)
        (inverse temp. phase transition properties of biodegradable)
RN
     223114-10-1 HCAPLUS
IT
     223114-10-1
     RL: PRP (Properties)
        (inverse temp. phase transition properties of biodegradable)
RE.CNT
RE
(1) Annaka, M; Nature 1992, V355, P430 HCAPLUS
(3) Chen, G; Nature 1995, V373, P49 HCAPLUS
(4) Cheng, Y; Macromolecules 1995, V28(8), P2665 HCAPLUS
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(5) Fujishige, S; J Phys Chem 1989, V93, P3311 HCAPLUS(6) Hirotsu, S; J Chem Phys 1987, V87(2), P1392 HCAPLUS

### hsu - 09 / 291426 ALL CITATIONS AVAILABLE IN THE RE FORMAT ANSWER 12 OF 29 HCAPLUS COPYRIGHT 2001 ACS 1999:109957 HCAPLUS DN 130:297251 Tyrosine-PEG-derived poly(ether carbonate)s as new biomaterials part. I: ΤI synthesis and evaluation ΑU Yu, Chun; Kohn, Joachim Department of Chemistry, Rutgers, The State University of New Jersey, New CS Brunswick, NJ, 08903, USA SO Biomaterials (1999), 20(3), 253-264 CODEN: BIMADU; ISSN: 0142-9612 PB Elsevier Science Ltd. DT Journal LA English Tyrosine-PEG-derived poly(ether carbonates) were prepd. by condensation AB copolymn. of PEG and deaminotyrosyltyrosine alkyl esters with phosgene. The resulting polymers were random copolymers with wt.-av. mol. wts. 40,000-200,000. Chem. structure and purity were confirmed by NMR and FTIR spectral anal. General structure-property correlations were established. The glass transition temp. decreased with increasing PEG content and increasing alkyl ester chain length. When higher mol. wt. PEG blocks were used, the glass transition temp. increased relative to identical polymers having shorter PEG blocks. The tensile modulus increased with decreasing PEG content, decreasing pendent chain length, and when longer PEG blocks were used. Water uptake and the rate of backbone degrdn. increased with increasing PEG content. Microspheres could be prepd. by solvent evapn. techniques from copolymers with low PEG content. Release rate of p-nitroaniline and fluorescein isothiocyanate-dextran from the microspheres increased with increasing PEG content. While tyrosine-derived polycarbonates were excellent substrates for cell attachment and growth, the presence of only 5 mol% of PEG1000 led to low or no cell attachment in short-term cell culture with both rat lung fibroblasts and osteoblasts. The polymers were non-cytotoxic. ΙT 223114-10-1P RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. and properties of tyrosine-based polyoxyethylenepolycarbonates) RN 223114-10-1 HCAPLUS ΙT 223114-10-1P 223114-11-2P 223114-13-4P 223114-15-6P RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. and properties of tyrosine-based polyoxyethylenepolycarbonates) RE.CNT 40 (1) Bakker, D; J Biomed Mater Res 1990, V24(4), P489 HCAPLUS (5) Brocchini, S; J Amer Chem Soc 1997, V119(19), P4553 HCAPLUS (6) Cho, C; J Biomed Mater Res 1993, V27, P199 HCAPLUS (7) Choueka, J; J Biomed Mater Res 1996, V31, P35 HCAPLUS (9) Engelberg, I; Biomaterials 1991, V12(3), P292 HCAPLUS ALL CITATIONS AVAILABLE IN THE RE FORMAT L93 ANSWER 13 OF 29 HCAPLUS COPYRIGHT 2001 ACS AN1998:531630 HCAPLUS DN 129:293785 Biosmart tyrosine polycarbonates ΤI ΑU Brode, G. L.; Kohn, J.; Kemnitzer, J. E. CS Integra LifeSciences Corp., Plainsboro, NJ, 08536, USA

Polym. Prepr. (Am. Chem. Soc., Div. Polym. Chem.) (1998), 39(2), 230-231

American Chemical Society, Division of Polymer Chemistry

SO

PB

DΤ

LA

Journal

English

CODEN: ACPPAY; ISSN: 0032-3934

AB Poly(desaminotyrosyltyrosine-co-desaminotyrosyltyrosine carbonate) was prepd. and was sol. at pH.gtoreq.6.0 in aq. or ionic buffer. Relevant medical applications are post-surgical antiadhesion barriers and oral drug delivery.

IT 214259-59-3DP, hydrogenated

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (Biosmart tyrosine polycarbonates)

RN 214259-59-3 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, polymer with carbonic acid and N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine ethyl ester (9CI) (CA INDEX NAME)

CM 1

CRN 135313-59-6 CMF C20 H23 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 86432-31-7 CMF C18 H19 N O5 CDES 5:L

## Absolute stereochemistry.

CM 3

CRN 463-79-6 CMF C H2 O3

IT 214259-59-3DP, hydrogenated

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (Biosmart tyrosine polycarbonates)

- L93 ANSWER 14 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 1997:738720 HCAPLUS
- DN 128:26780
- TI Amino acid derived **polymers** for use in controlled delivery systems of peptides
- AU Brocchini, S.; Schachter, D. M.; Kohn, J.
- CS Department of Chemistry, Rutgers, The State University of New Jersey, New Brunswick, NJ, 08903, USA
- SO ACS Symp. Ser. (1997), 675 (Therapeutic Protein and Peptide Formulation and Delivery), 154-167
  CODEN: ACSMC8; ISSN: 0097-6156
- PB American Chemical Society
- DT Journal; General Review
- LA English
- AB A review with 32 refs. A family of synthetic, tyrosine-derived polyarylates is being studied as a new polymeric matrix system for the controlled release of peptides. The polyarylates are degradable amorphous materials whose backbone structure contains amide bonds. Using the cyclic heptapeptide contained in the platelet integrin glycoprotein IIb/IIIa blocking formulation INTEGRILIN as a model, the effect of the polymer structure on peptide miscibility within the polymeric matrix and its effect on the release behavior was investigated. A new co-pptn. technique provided polyarylate-peptide blends that were compression molded without decompn., deactivation, or detectable aggregation of the peptide. Transparent, pliable compression molded films with high peptide loadings of up to 50% (wt./wt.) were fabricated in this way. In spite of such high loadings, the model peptide was not released from these films over a 30 day exposure to physiol. buffer soln. at 37 .degree.C. Only when polyethylene glycol (PEG) was added to the formulation was the model peptide released. Release rate was a function of polyarylate structure and the amt. and mol. wt. of PEG used in the blends. This provided an effective means to modulate the release rate.
- L93 ANSWER 15 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 1997:336802 HCAPLUS
- DN 127:8987
- TI Pseudo-poly(amino acid)s. Examples for synthetic materials derived from natural metabolites
- AU James, Kenneth; Kohn, Joachim
- CS Department of Chemistry, Rutgers The State University of New Jersey, New Brunswick, NJ, 08903, USA
- SO Controlled Drug Delivery (1997), 389-403. Editor(s): Park, Kinam. Publisher: American Chemical Society, Washington, D. C. CODEN: 64KFAX
- DT Conference; General Review
- LA English
- AB A review with 57 refs. Pseudo-poly(amino acid)s consisting of .alpha.-L-amino acid building blocks linked by nonamide bonds have been used successfully as carriers in direct intracranial drug delivery, immunizations systems, and the controlled release of anticoagulants. These natural metabolite-based polymers are biocompatible, degradable materials readily processed into microspheres, films, fibers, pins, and screws. In the last 5 yr, most attention has been directed toward tyrosine-derived polycarbonates, polyiminocarbonates, and polyarylates. Pseudo-poly(aminoacid) chem. and synthesis, physicomech. and degrdn. properties, biol. response, immunol. considerations, sterilization, and drug delivery applications thus far investigated are summarized.
- L93 ANSWER 16 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 1997:230323 HCAPLUS
- TI A combinatorial approach for the development of new polymer biomaterials.
- AU Brocchini, S.; James, K. S.; Tangpasuthadol,

V.; Kohn, J.

CS Department Chemistry, Rutgers University, Piscataway, NJ, 08855, USA

- SO Book of Abstracts, 213th ACS National Meeting, San Francisco, April 13-17 (1997), BIOC-257 Publisher: American Chemical Society, Washington, D. C. CODEN: 64AOAA
- DT Conference; Meeting Abstract
- LA English
- Development of synthetic degradable polymers for use in medical AΒ applications including tissue replacement therapy is a complex challenge. These medical polymers must meet a multitude of stringent requirements similar in kind to drug candidates while also exhibiting appropriate physicomech. properties that meet the criteria for a given application. Of central importance is the requirement to understand the influence of polymer structure on the cellular response. report the concept of creating a permutationally designed polymer system to prep. a library of polymers that can be used (a) to increase the no. of available polymer candidate materials for medical applications, and (b) to systematize the study of correlations between polymer structure, material properties, and biol. responses. In this combinatorial approach, structural variations of a small no. of monomers were systematically translated into a library of 112 tyrosinederived polyarylates which were prepd. on a 0.2 g scale in arrays of glass vials set up in a shaker bath. Up to 32 simultaneous reactions were conducted in a run and the polyarylates were characterized by mol. wt., Tg, and air-water contact angle. Properties, including in vitro fibroblast proliferation, incrementally varied over a wide range. This resulted in several structure-property correlations which will be described.
- L93 ANSWER 17 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 1997:36235 HCAPLUS
- DN 126:65248
- TI New biomaterials for tissue engineering
- AU James, Kenneth; Kohn, Joachim
- CS USA
- SO MRS Bull. (1996), 21(11), 22-26 CODEN: MRSBEA; ISSN: 0883-7694
- PB Materials Research Society
- DT Journal; General Review
- LA English
- AB A review with 41 refs., esp. on tyrosine-derived polycarbonates and polyarylates.
- L93 ANSWER 18 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 1997:14265 HCAPLUS
- TI Applications of pseudo-poly(amino acid) biomaterials
- AU James, Kenneth; Kohn, Joachim
- CS Dep. Chem., Rutgers Univ., Piscataway, NJ, 08855-0939, USA
- SO Trends Polym. Sci. (Cambridge, U. K.) (1996), 4(12), 394-397 CODEN: TPSCE8; ISSN: 0966-4793
- PB Elsevier
- DT Journal; News Announcement
- LA English
- AB Unavailable
- L93 ANSWER 19 OF 29 HCAPLUS COPYRIGHT 2001 ACS
- AN 1996:250260 HCAPLUS
- DN 124:325298
- TI Canine bone response to tyrosine-derived polycarbonates and poly(L-lactic acid)
- AU Choueka, Jack; Charvet, Jose L.; Koval, Kenneth J.; Alexander, Harold; James, Kenneth S.; Hooper, Kimberly A.; Kohn, Joachim
- CS Dep. Bioengineering, Orthopaedic Inst., New York, NY, USA
- SO J. Biomed. Mater. Res. (1996), 31(1), 35-41
- CODEN: JBMRBG; ISSN: 0021-9304
- DT Journal

LA English AΒ Tyrosine-derived polycarbonates are a new class of degradable polymers developed for orthopedic applications. this study the long-term (48 wk) in vivo degrdn. kinetics and host bone response to poly(DTE carbonate) and poly(DTH carbonate) were investigated using a canine bone chamber model. Poly(L-lactic acid) (PLA) served as a control material. Two chambers of each test material were retrieved at 6-, 12-, 24-, and 48-wk time points. Tyrosine-derived polycarbonates were found to exhibit degrdn. kinetics comparable to PLA. Each test material lost approx. 50% of its initial mol. wt. (Mw) over the 48-wk test period. Poly(DTE carbonate) and poly(DTH carbonate) test chambers were characterized by sustained bone ingrowth throughout the 48 wk. In contrast, bone ingrowth into the PLA chambers peaked at 24 wk and dropped by half at the 48-wk time point. A fibrous tissue layer was found surrounding the PLA implants at all time points. This fibrous tissue layer was notably absent at the interface between bone and the tyrosine-derived polycarbonates. Histol. sections revealed intimate contact between bone and tyrosine-derived polycarbonates. From a degrdn.-biocompatibility perspective, the tyrosine-derived polycarbonates appear to be comparable, if not superior, to PLA in this canine bone chamber model. L93 ANSWER 20 OF 29 HCAPLUS COPYRIGHT 2001 ACS AN 1993:450276 HCAPLUS DN 119:50276 ΤI Physicomechanical properties of biodegradable polymers for medical applications ΑU Engelberg, I.; Kohn, J. CS RAFAEL - Armament Dev. Author., Haifa, 31021, Israel SO Isr. Mater. Eng. Conf., 5th (1991) 277-91 CODEN: 58SIAV DT Journal LA English AΒ The physicomech. properties of degradable polymers used for medical applications were characterized. The following polymers were included in this study: three samples of poly(ortho esters) derived from 3,9-bis(ethylidene-2,4,8,10-tetraoxaspiro[5.5]undecane) and various ratios of 1,6-hexanediol and trans-cyclohexanedimethanol, poly(glycolic acid), six samples of poly(L-lactic acid) and poly(D,L-lactic acid) with mol. wts. 21,000-550,000 dalton, poly(.epsilon.-caprolactone), poly(.beta.-hydroxybutyrate) and three copolymers of .beta.-hydroxybutyric acid and various amts. of hydroxyvaleric acid, one sample each of two different types of poly(anhydrides), poly(trimethylene carbonate), and two different poly(iminocarbonates). For each polymer, the thermal properties (glass transition temp., crystn., melting and decompn. points) were detd. by DSC and by TGA. The tensile properties (Young's modulus, tensile strength, and elongation at yield and break) were detd. by tensile testing on an Instron stress-strain tester. The flexural storage modulus as a function of temp. was detd. by dynamic mech. anal. ΙT 133063-34-0 RL: PRP (Properties) (biodegradable, physicomech. properties of, for medical applications) RN 133063-34-0 HCAPLUS CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, homopolymer (9CI) (CA INDEX NAME) CM 1

Absolute stereochemistry.

CDES 5:L

CRN 133063-33-9 CMF C24 H31 N O5

IT 133063-34-0

RL: PRP (Properties)

(biodegradable, physicomech. properties of, for medical applications)

L93 ANSWER 21 OF 29 HCAPLUS COPYRIGHT 2001 ACS

1992:455888 HCAPLUS ΑN

DN 117:55888

TI Tyrosine-derived polycarbonates: backbone-modified "pseudo"-poly(amino acids) designed for biomedical applications

ΑU Pulapura, Satish; Kohn, Joachim

Dep. Chem., Rutgers, State Univ., New Brunswick, NJ, 08855-0939, USA Biopolymers (1992), 32(4), 411-17 CS

SO CODEN: BIPMAA; ISSN: 0006-3525

DT Journal

LA English

AΒ

Starting from L-tyrosine (Tyr) and its metabolites desaminotyrosine (Dat) and tyramine (Tym), four structurally related model dipeptides were prepd.: Dat-Tym (neither N- or C-terminus present), Z-Tyr-Tym (N-terminus protected by benzyloxycarbonyl) Dat-Tyr-Hex (C-terminus protected by a hexyl ester group), and Z-Tyr-Tyr-Hex (both N- and C-termini present, protected by benzyloxycarbonyl and hexyl ester, resp.). The model dipeptides were used a monomers in the synthesis of polycarbonates. The polymn. reaction in the presence of either phosgene or triphosgene proceeded via the phenolic hydroxyl groups. Polymers with mol. wts. of 105,000-400,000 Datlon (by gel permeation chromatog., relative to polystyrene stds.) were obtained. The physicomech. properties (soly., mech. strength, glass transition and decompn. temp., processibility) of the polymers were detd., and an attempt was made to correlate the polymer properties with the nature of the N- and C-terminus protecting groups. The presence of the urethane bond at the N-terminus protecting group was found to reduce soly., ductility, and processibility, probably due to interchain hydrogen bonding. The presence of a C-terminus alkyl ester group increased soly. and processibility. Thus, the most promising candidate polymer for biomedical applications was obtained from Dat-Tyr-Hex, the monomer carrying a C-terminus protecting group only. Since very similar results had recently been obtained for a series of structurally related polyiminocarbonates, the structure property correlations seems to be generally valid.

IT

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and physicomech. properties of, structure effect on, biomaterials in relation to)

RN 133063-34-0 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

### IT 133063-34-0P 133063-35-1P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and physicomech. properties of, structure effect on, biomaterials in relation to)

L93 ANSWER 22 OF 29 HCAPLUS COPYRIGHT 2001 ACS

AN 1990:578192 HCAPLUS

DN 113:178192

TI Biomaterials based on "pseudo"-poly(amino acids): a study of tyrosine-derived polyiminocarbonates

AU Pulapura, S.; Kohn, J.

CS Dep. Chem., Rutgers Univ., New Brunswick, NJ, 08855, USA

SO Polym. Prepr. (Am. Chem. Soc., Div. Polym. Chem.) (1990), 31(1), 233-4 CODEN: ACPPAY; ISSN: 0032-3934

DT Journal

LA English

GI

AB Polyiminocarbonates, e.g. I (R = Et, hexyl, or palmityl) were prepd. and their properties detd. Incorporation of nonamide linkages into the backbone of the poly(amino acids) leads to an improvement of the processibility and the physicomech. properties of the polymers. None of the polymers exhibited gross toxicity or tissue incompatibility on s.c. implantation in mice, rats, or rabbits.

IT 106231-87-2P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and properties of, for biomaterials)

RN 106231-87-2 HCAPLUS

CN L-Tyrosine, N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, 4-carbamate, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 106231-86-1 CMF C33 H39 N3 O8 CDES 5:L,L

### IT 106231-87-2P 128835-54-1P 128835-60-9P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation) (prepn. and properties of, for biomaterials)

L93 ANSWER 23 OF 29 HCAPLUS COPYRIGHT 2001 ACS

AN 1990:119400 HCAPLUS

DN 112:119400

TI Backbone modification of synthetic poly-.alpha.-L-amino acids

AU Kohn, Joachim; Langer, Robert

CS Dep. Chem., Rutgers, State Univ., Piscataway, NJ, 08855-0939, USA

SO Pept.: Chem. Biol., Proc. Am. Pept. Symp. 10th (1988), Meeting Date 1987, 658-61. Editor(s): Marshall, Garland R. Publisher: ESCOM Sci. Pub., Leiden, Neth. CODEN: 56MDA6

DT Conference

LA English

GΙ

AB A symposium on the prepn. of tyrosyltyrosine side chain polymers I (R = Et, hexyl, palmityl). The polymers show promise as biodegradable implantable drug delivery devices.

IT 93174-02-8P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn., melting range, and soly. of, in methylene chloride)

RN 93174-02-8 HCAPLUS

CN L-Tyrosine, N-[O-cyano-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, ethyl ester, cyanate (ester), polymer with N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-L-tyrosine ethyl ester (9CI) (CA INDEX NAME)

CM 1

CRN 93174-01-7 CMF C30 H28 N4 O7 CDES 5:L,L

CM 2

CRN 4142-95-4 CMF C28 H30 N2 O7 CDES 5:L,L

Absolute stereochemistry.

# IT 93174-02-8P 118949-20-5P 125607-28-5P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn., melting range, and soly. of, in methylene chloride)

L93 ANSWER 24 OF 29 HCAPLUS COPYRIGHT 2001 ACS

AN 1989:595409 HCAPLUS

DN 111:195409

TI Preparation of nonpeptide polyamino acid bioerodible polymers for drug formulations

IN Kohn, Joachim; Langer, Robert S.

PA Massachusetts Institute of Technology, USA

SO U.S., 9 pp. CODEN: USXXAM

DT Patent

LA English

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	US 4638045	Α	19870120	US 1985-703153	19850219
	US 4863735	A	19890905	US 1986-914380	19861002
DDAT	IIC 1005_702152	10050	210		

Biodegradable polymers are prepd. by polymn. of ZNHCHR1CONHCHR2COY or ZNHCHR1CONHCHR2COYHCHR3COY (R1-R3 = side chains of L-.alpha.-amino acids; Y, Z = protecting group) through .gtoreq.l of R1-R3, useful for controlled release applications in vivo and vitro for delivery of a wide variety of biol. and pharmacol. active ligands. are prepd. Thus, Z-Glu-Phe-OH (Z = PhCH2O2C) and Et3N in CH2Cl2 were treated with (PhO)2P(O)Cl and the mixt. was kept at 4.degree. for 12 h to give a polymer with an av. mol. wt. of 17,000.

IT 123375-14-4P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of, as bioerodible material)

RN 123375-14-4 HCAPLUS

CN L-Tyrosine, N-[O-(aminocarbonyl)-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, ethyl ester, carbamate (ester), homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 123375-13-3 CMF C30 H32 N4 O9 CDES 5:L,L

Absolute stereochemistry.

IT 123375-14-4P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of, as bioerodible material)

L93 ANSWER 25 OF 29 HCAPLUS COPYRIGHT 2001 ACS

AN 1989:213501 HCAPLUS

DN 110:213501

TI Synthesis of poly(iminocarbonates): degradable polymers with potential applications as disposable plastics and as biomaterials

AU Li, Chun; Kohn, Joachim

CS Dep. Chem., Rutgers, State Univ., New Brunswick, NJ, 08903, USA

SO Macromolecules (1989), 22(5), 2029-36

CODEN: MAMOBX; ISSN: 0024-9297

DT Journal

LA English

AΒ The prepn. of poly(iminocarbonates) by bulk, soln., and interfacial polycondensation was studied by model copolymn. of bisphenol A (I) with bisphenol A dicyanate (II). Bulk polymn. was unsuitable for prepn. of structurally well-defined poly(iminocarbonates) due to the formation of crosslinked products. The soln. polymn. of I with II was optimized in terms of solvent, catalyst, and catalyst concn. In THF with K tert-butoxide as the catalyst, polymers with mol. wts. up to 80,000 were obtained. Interfacial polymn. was a feasible prepn. technique in spite of the high sensitivity of cyanates toward hydrolysis. In the presence of Bu4NBr as a phase-transfer catalyst, polymers with mol. wts. >100,000 were obtained. The interfacial polymn. of I with CNBr instead with II was also feasible, eliminating the need to isolate and purify the reactive dicyanate. Novel poly(iminocarbonates) were prepd. by soln. polymn. of II with 4,4'-thiodiphenol, desaminotyrosyl tryramine, or N-(benzoyloxy)carbonyl-L-tyrosyl-L-tyrosine hexyl ether. The potential applications of poly(iminocarbonates) as disposable plastics and biomaterials were discussed.

RN 118798-95-1 HCAPLUS

CN L-Tyrosine, N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester,
 polymer with (1-methylethylidene)di-4,1-phenylene dicyanate (9CI) (CA
 INDEX NAME)

CRN 106231-84-9 CMF C32 H38 N2 O7 CDES 5:L,L

Absolute stereochemistry.

CM 2

CRN 1156-51-0 CMF C17 H14 N2 O2

IT 118798-95-1P, N-(Benzyloxy)carbonyl-L-tyrosyl-L-tyrosine hexyl ester-bisphenol A dicyanate copolymer

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and characterization of)

L93 ANSWER 26 OF 29 HCAPLUS COPYRIGHT 2001 ACS

AN 1989:101818 HCAPLUS

DN 110:101818

TI Implantable biodegradable polymeric drug delivery system with adjuvant activity

IN Kohn, Joachim B.; Langer, Robert S.; Niemi, Steven M.; Fox, James G.

PA Massachusetts Institute of Technology, USA

SO PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN. CNT 2

CAN.	CNT Z			
	PATENT NO.	KIND DATE	APPLICATION NO.	DATE
PI	WO 8802262 W: JP	A1 19880407	WO 1987-US2428	19870924
	RW: AT, BE, US 4863735	CH, DE, FR, GB, IT, A 19890905	US 1986-914380	19861002
	CA 1323566	A1 19931026	CA 1987-548151	19870929
PRAI	US 1986-914380	19861002		
	US 1985-703153	19850219		

AB An antigen delivery system is described which utilizes a biodegradable polymer with good mech. properties in combination with a material stimulating the immune system. The material having adjuvant activity maybe a polymer degrdn. product or an adjuvant which is contained within

or bound to the polymer. The polymer may be formed from tyrosine dipeptides. This polymer is not an adjuvant, but degrades into products which stimulate the immune system. The advantages of this system are that a polymer can be used to form a biodegradable integral structure which is useful as both an implantable source of an antigen or other biol. active compds. and as a control means for the rate of release of the biol. active compd., producing a sustained, relatively const. delivery of antigen with simultaneous stimulation of the immune response. N-Cbz-Tyr-Tyr-Hex (CTTH; Cbz = benzyloxycarbonyl; Hex = hexyl) (prepn. given) was treated with CNBr to form the dicyanate, which was mixed with an equimolar quantity of CTTH and the mixt. polymd. in the presence of KOBu-tert to give poly(CTTH-iminocarbonate) with mol. wt. 19,500 and which had an intrinsic viscosity 0.27 (DMF, 25.degree.). Using particulate suspensions, the degrdn. products of poly(CTTH-iminocarbonate) were shown to be as potent an adjuvant as complete Freund's adjuvant and muramyl dipeptide when the serum antibody response to bovine serum albumin (BSA) in male CD-1 mice is measured over 56 wk. Further, BSA released from s.c. implanted polymeric antigen delivery devices made of poly(CTTH-iminocarbonate) resulted in significantly higher anti-BSA titers than devices made of poly(bisphenol A-iminicarbonate). At the end of week 56 the injection and implantation sites were examd. and no tissue abnormalities were found, no residues of the injected particulate material were detected, and only small amts. of polymeric residue were detected at the implantation sites.

IT 118949-20-5P

RL: PREP (Preparation)

(manuf. of, as biodegradable polymer for stimulation of immune response)

RN 118949-20-5 HCAPLUS

L-Tyrosine, N-[O-cyano-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, cyanate (ester), polymer with N-[N-{(phenylmethoxy)carbonyl]-L-tyrosyl]-L-tyrosine hexyl ester (9CI) (CA INDEX NAME)

CM 1

CN

CRN 106231-84-9 CMF C32 H38 N2 O7 CDES\_5:L,L

Absolute stereochemistry.

CM 2

CRN 104549-44-2 CMF C34 H36 N4 O7 CDES 5:L,L

#### IT 118949-20-5P

RL: PREP (Preparation)

(manuf. of, as biodegradable polymer for stimulation of immune response)

L93 ANSWER 27 OF 29 HCAPLUS COPYRIGHT 2001 ACS

AN 1987:162477 HCAPLUS

DN 106:162477

TI Single-step immunization using a controlled release, biodegradable polymer with sustained adjuvant activity

AU Kohn, Joachim; Niemi, Steven M.; Albert, Elizabeth C.; Murphy,

James C.; Langer, Robert; Fox, James G.

CS Dep. Appl. Biol. Sci., Massachusetts Inst. Technol., Cambridge, MA, 02139, USA

SO J. Immunol. Methods (1986), 95(1), 31-8

CODEN: JIMMBG; ISSN: 0022-1759

DT Journal

LA English

GI

The use of a biodegradable polymer for antigen delivery based on poly(CTTH-iminocarbonate) (I) [106755-33-3] was investigated. This polymer was selected since its primary degrdn. product, N-benzyloxycarbonyl-L-tyrosyl-L-tyrosine hexyl ester (CTTH) [106231-84-9] was as potent an adjuvant as complete Freund's adjuvant and muramyl dipeptide, when measuring the serum antibody response to bovine serum albumin (BSA) in male CD-1 mice over 56 wk. BSA released from s.c. implanted polymeric antigen delivery devices made of I resulted in significantly higher anti-BSA antibody titers than devices made of poly(bisphenol A-iminocarbonate) [26101-32-6].

IT 107173-10-4

RL: BIOL (Biological study)

(biodegradable, with sustained adjuvant activity)

RN 107173-10-4 HCAPLUS

CN L-Tyrosine, N-[O-cyano-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, cyanate (ester), homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 104549-44-2

CMF C34 H36 N4 O7 CDES 5:L,L

Absolute stereochemistry.

IT 107173-10-4

RL: BIOL (Biological study)

(biodegradable, with sustained adjuvant activity)

L93 ANSWER 28 OF 29 HCAPLUS COPYRIGHT 2001 ACS

AN 1987:67663 HCAPLUS

DN 106:67663

TI Polymerization reactions involving the side chains of .alpha.-L-amino acids

AU Kohn, Joachim; Langer, Robert

CS Dep. Appl. Biol. Sci., Massachusetts Inst. Technol., Cambridge, MA, 02139, USA

SO J. Am. Chem. Soc. (1987), 109(3), 817-20

CODEN: JACSAT; ISSN: 0002-7863

DT Journal

LA English

OS CASREACT 106:67663

GI

AB Hydroxyproline deriv. I (Pal = palmitoyl) underwent melt transesterification in the presence of Al isopropoxide to give side-chain polymer II. Z-Tyr-Tyr-OHex (III; Z = PhCH2O2C, Hex = hexyl) was treated with cyanogen bromide to give dicyanate IV. The soln. polymn. of equimolar amts. of III and IV in THF contg. KOCMe3 gave the corresponding iminocarbonate side-chain polymer.

IT 106231-87-2P

RN 106231-87-2 HCAPLUS

CN L-Tyrosine, N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, 4-carbamate, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 106231-86-1 CMF C33 H39 N3 O8 CDES 5:L,L

Absolute stereochemistry.

IT 106231-87-2P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of)

L93 ANSWER 29 OF 29 HCAPLUS COPYRIGHT 2001 ACS

AN 1984:616333 HCAPLUS

DN 101:216333

A new approach to the development of bioerodible polymers for controlled ΤI release applications employing naturally occurring amino acids

ΑU

Kohn, Joachim; Langer, Robert
Whitaker Coll. Health Sci., Massachusetts Inst. Technol., Cambridge, MA, CS 02139, USA

Polym. Mater. Sci. Eng. (1984), 51, 119-21 SO

CODEN: PMSEDG

DT Journal

LA English

AB Dipeptides, e.g., N-carbobenzoxytyrosyltyrosine Et ester (I) [4142-95-4], can be used as monomers for the formation of nonpeptide polymers for controlled-release of drugs. I was prepd. by known methods having 2 reactive, arom. OH groups which could be used for formation of a hydrolytically labile iminocarbonate linkage. I-di-O-cyano-I copolymer [ 93174-02-8] erodes completely within 93 days when exposed to 0.1M phosphate buffer (pH 7.4) at 37.degree..

ΙT 93174-02-8P

> RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and biodegradability of, controlled-release applications in relation to)

RN 93174-02-8 HCAPLUS

CN L-Tyrosine, N-[O-cyano-N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, ethyl ester, cyanate (ester), polymer with N-[N-[(phenylmethoxy)carbonyl]-Ltyrosyl]-L-tyrosine ethyl ester (9CI) (CA INDEX NAME)

CM

CRN 93174-01-7 CMF C30 H28 N4 O7 CDES 5:L,L

CM 2

CRN 4142-95-4 CMF C28 H30 N2 O7 CDES 5:L,L

Absolute stereochemistry.

IT 93174-02-8P

RL: SPN (Synthetic preparation); PREP (Preparation)
(prepn. and biodegradability of, controlled-release applications in relation to)

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L101 ANSWER 1 OF 20 HCAPLUS COPYRIGHT 2001 ACS
     1998:706136 HCAPLUS
ΑN
DN
     129:321238
ΤI
     Biodegradable and biocompatible polymer for use as medical implant and
     drug-delivery system.
     Mao, Hai-quan; Leong, Kam W.
IN
PΑ
     Johns Hopkins University School of Medicine, USA
SO
     PCT Int. Appl., 55 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                      KIND
                             DATE
                                            APPLICATION NO.
                                                              DATE
PΙ
     WO 9846286
                       A1
                             19981022
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PATENT NO. KIND DATE APPLICATION NO. DATE

WO 9846286 Al 19981022 WO 1998-US7585 19980414 <-W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG

US 5912225 A 19990615 US 1997-834164 19970414 <--

AU 1998-71206

AU 9871206 A1 19981111 PRAI US 1997-834164 19970414 <--WO 1998-US7585 19980414 19980414 <--

GI

AB The title polymers are I (R = H, alkyl, aryl or heterocyclyl; R1 = R, alkoxy, aryloxy or heterocyclyloxy; n = 5 to 500). They are biodegradable and biocompatible before and upon biodegrdn. Processes for prepg. the polymers, compns. contg. the polymers and biol. active substances, articles useful for implantation or injection into the body fabricated from the compns., and methods for controllably releasing biol. active substances using the polymers, are also described.

IT 214957-41-2P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. as biodegradable and biocompatible polymer)

RN 214957-41-2 HCAPLUS

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with ethyl phosphorodichloridate (9CI) (CA INDEX NAME)

CM· 1

CRN 133063-33-9 CMF C24 H31 N O5 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 1498-51-7 CMF C2 H5 C12 O2 P

#### IT 214957-41-2P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. as biodegradable and biocompatible polymer)

L101 ANSWER 2 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:458330 HCAPLUS

DN 127:166621

TI Syntheses of poly(phosphate-urethane)s as drug release materials

AU Zhuro, Ren-Xi; Wang, Jun; Mao, Hai-Quan

CS Dep. Chem., Laboratory Biomedical Polymer Materials State Education Commission China, Wuhan University, Wuhan, 430072, Peop. Rep. China SO Gaodeng Xuexiao Huaxue Xuebao (1997), 18(7), 1207-1211

Gaodeng Xuexiao Huaxue Xuebao (1997), 18(7), 1207-1211 CODEN: KTHPDM; ISSN: 0251-0790

PB Gaodeng Jiaoyu Chubanshe

DT Journal

LA Chinese

AB New biodegradable and biocompatible copolymers poly(phosphate-urethane)s with elevated mol. wt. were synthesized by the polycondensation of Ser-Tyr or Tyr-Tyr dipeptide and phosphorodichloridate and followed by reacting with 1,4-butane diisocyanate. The chem. structures were confirmed by 1H NMR, FTIR and elemental anal. The mol. wt., and thermostability of these polymers were investigated. The in vitro degrdn. and bovine serum albumin (BSA) release profile were also studied. It was found that the degrdn. rate of the materials and BSA release behavior can be modulated by varying the hydrohobic properties of these poly(phosphate-urethane)s.

IT 193526-44-2P

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. of poly(phosphate-urethane)s as drug release materials)

RN 193526-44-2 HCAPLUS

CN L-Tyrosine, N-[(phenylmethoxy)carbonyl]-L-tyrosyl-, dodecyl ester, polymer
with 1,4-diisocyanatobutane and phenyl phosphorodichloridate (9CI) (CA
INDEX NAME)

CM 1

CRN 193526-43-1 CMF C38 H50 N2 O7

### Absolute stereochemistry.

CM 2

CRN 4538-37-8 CMF C6 H8 N2 O2

OCN-(CH<sub>2</sub>)<sub>4</sub>-NCO

CM 3

CRN 770-12-7

CMF C6 H5 C12 O2 P

WO 1995-US9614

GI

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Cl-P-OPh
   Cl
IT
     193526-44-2P 193526-45-3P
     RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (prepn. of poly(phosphate-urethane)s as drug release materials)
L101 ANSWER 3 OF 20 HCAPLUS COPYRIGHT 2001 ACS
ΑN
     1996:369153 HCAPLUS
DN
     125:34037
ΤI
     Preparation of soluble combinatorial libraries using soluble
     macromolecular supports
     Janda, Kim; Han, Hyunsoo
IN
PΑ
     Scripps Research Institute, USA
     PCT Int. Appl., 154 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                      KIND
                           DATE
                                          APPLICATION NO. DATE
                                          -----
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ΡI
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                      A1
                           19960208
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             MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ,
             TM, TT
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             LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE,
                    ΤG
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                                          AU 1995-32722
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                            19981022
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                                          EP 1995-929334
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                      T2 19980623
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PRAI US 1994-281200
                     19940726 <--
    US 1995-484153
                     19950607 <--
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# \* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

19950726 <--

AB Novel sol. combinatorial libraries are prepd., comprising a sol. phase in soln. attached to a core mol., and allowing the improved high-yield and efficient prodn. of sol. combinatorial libraries. Some specific examples of the sol. combinatorial libraries claimed herein comprise one or more of the following: amino acids, .alpha.-azetide amino acids, triazine dione mols., .gamma.-lactamtide mols. (constrained peptide mimics), .delta.-lactamthiotide mols. (constrained peptide mimics), .beta.-lactam nucleus contg. mols., lycoramine alkaloid nucleus contg. mols., .beta.-blocker nucleus mols., oligopeptides, oligosaccharides, oligonucleotides, and arylsulfonamides. The macromol. supports are selected from polyethylene glycol, polyvinyl alc., polyvinylamine copolymd. with polyvinylpyrrolidine, and derivs. thereof. Further, a split synthesis technique for generating libraries of combinatorial mols.

employs a biphasic macromol. support which is sol. during the pooling, splitting, and coupling steps but which is insol. during the washing step. The use of a biphasic macromol. support in its insol. phase significantly enhances the efficiency and performance of the washing step. Thus, a library of 8 tetrasaccharides (e.g. I, II, and III), useful as antigenic markers which distinguishes fetal erythrocytes from adult cells (no data), were prepd. by the split synthesis technique involving sequential coupling of a library of polyethylene glycol monomethyl ether-bound glucose and galactose derivs. (IV and V; R = MeO-PEG-O2CCH2CH2CO, wherein PEG = polyethylene glycol) (prepn. given) with (A) galactosamine and glucosamine derivs. (VI and VII) (prepn. given), (B) glucose and galactose derivs. IV and V (R = H) (prepn. given), and (C) galactosamine deriv. VI.

IT 169692-78-8P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(antigenic determinant recognized by monoclonal antibody 3ET; prepn. of sol. combinatorial libraries using sol. macromol. supports)

RN 169692-78-8 HCAPLUS

CN Poly(oxy-1,2-ethanediyl), .alpha.-methyl-.omega.-hydroxy-, ester with N-[N-[N-(N-L-tyrosylglycyl)glycyl]-L-phenylalanyl]-L-tyrosine (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-B

#### IT 169692-78-8P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(antigenic determinant recognized by monoclonal antibody 3ET; prepn. of sol. combinatorial libraries using sol. macromol. supports)

L101 ANSWER 4 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1995:676982 HCAPLUS

DN 123:286659

TI Liquid-phase combinatorial synthesis

AU Han, Hyunsoo; Wolfe, Mary M.; Brenner, Sydney; Janda, Kim D.

CS Dep. Mol. Biol. Chem., Scripps Res. Inst., La Jolla, CA, 92037, USA

SO Proc. Natl. Acad. Sci. U. S. A. (1995), 92(14), 6419-23 CODEN: PNASA6; ISSN: 0027-8424

DT Journal

LA English

AB A concept termed liq.-phase combinatorial synthesis (LPCS) is described. The central feature of this methodol. is that it combines the advantages that classic org. synthesis in soln. offers with those that solid-phase synthesis can provide, through the application of a linear homogeneous polymer. To validate this concept two libraries were prepd., one of peptide and the second of nonpeptide origin. The peptide-based library was synthesized by a recursive deconvolution strategy (E. Erb, et al., 1994), and several ligands found in this library bind a monoclonal antibody elicited against .beta.-endorphin. The non-peptide mols. were arylsulfonamides, a class of compds. of known clin. bactericidal efficacy. The results indicate that the reaction scope of LPCS should be general, and its value to multiple, high-throughput screening assays could be of particular merit, since multi-milligram quantities of each library member can readily be attained.

IT 169692-78-8P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(liq.-phase combinatorial synthesis of peptides and arylsulfonamides)

RN 169692-78-8 HCAPLUS

CN Poly(oxy-1,2-ethanediyl), .alpha.-methyl-.omega.-hydroxy-, ester with N-[N-[N-(N-L-tyrosylglycyl)glycyl]-L-phenylalanyl]-L-tyrosine (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-B

# IT 169692-78-8P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation) (liq.-phase combinatorial synthesis of peptides and arylsulfonamides)

- L101 ANSWER 5 OF 20 HCAPLUS COPYRIGHT 2001 ACS
- AN 1995:294200 HCAPLUS
- DN 122:64325
- TI Drug-delivery polymers and pharmaceutical compositions employing them
- IN Kopecek, Jindrich; Rejmanova, Pavla; Strohalm, Jiri; et al.
- PA Ustav Makromolekularni Chemie AVCR, Czech Rep.
- SO Czech Rep., 50 pp.
- CODEN: CZXXED

Czech

DT Patent

LA

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE	
ΡI	CZ 278551	В6	19940316	CZ 1985-97	19850104	<
	SK 278506	В6	19970806	SK 1985-97	19850104	<
PRAI	CZ 1985-97	19850	104 <			

AB Drug-delivery polymers can be prepd. which are composed 5.0-99.7 mol% of units derived from Me-C:CH2-CO-NH-CH2-CHOH-Me, 0.2-20.0 mol% of units having the structure Me-C:CH2-CO-[NH-R-CO]-[B], where B is a bioactive mol. or drug, and 0.1-94.8 mol% of units having the structure Me-C:CH2-CO-NH-[D] or Me-C:CH2-CO-[D] or Me-C:CH2-CO-[NH-R-CO]-D, where D is a determinant and [NH-R-CO] is a spacer residue derived from Leu, Phe, Gly-Gly, Gly-Leu-Gly, Gly-Val-Ala, Gly-Phe-Ala, Gly-Leu-Phe, Gly-Leu-Ala, Ala-Val-Ala, Gly-Phe-Leu-Gly, Gly-Phe-Phe-Leu, Gly-Leu-Leuy-Gly, Gly-Phe-Tyr-Ala, Gly-Phe-Gly-Phe, Ala-Gly-Val-Phe, Gly-Phe-Phe-Gly, Gly-Phe-Leu-Gly-Phe, or Gly-Gly-Phe-Leu-Gly-Phe. Copolymers contg. the above components can be single or double-chained and may contain as bioactive mols. antitumor drugs, antimicrobials, parasiticides, antiinflammatories, cardiovascular agents, or nervous system agents. determinants may be monosaccharides, disaccharides, oligosaccharides, or O-methacryloylated sugars, which are preferably linked by an amide bond to an antibody such as IgG or anti-O antibody, or a protein such as transferrin, or a hormone such as MSH. Suitable determinants are galactose, galactosamine, glucosamine, mannosamine, and fucosylamine. The peptide spacers are degradable by lysosomal enzymes, releasing the pharmacol. active agents after the copolymer is taken up by target cells. Data are presented on the antileukemic activity of several claimed copolymers against leukemia L1210, and antitumor activity against melanoma and human hepatoma.

IT 79637-25-5DP, conjugates with bleomycin

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of drug-delivery polymers and pharmaceutical compns. employing them)

RN 79637-25-5 HCAPLUS

CN L-Tyrosine, N-[N-[N-(2-methyl-1-oxo-2-propenyl)glycyl]-L-phenylalanyl]-,
4-nitrophenyl ester, polymer with N-(2-hydroxypropyl)-2-methyl-2propenamide (9CI) (CA INDEX NAME)

CM 1

CRN 79637-24-4 CMF C30 H30 N4 O8 CDES 5:L,L

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CM 2
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CRN 21442-01-3 CMF C7 H13 N O2

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\begin{array}{c|cccc} \text{OH} & \text{O} & \text{CH}_2 \\ | & || & || \\ \text{Me-CH-CH}_2 - \text{NH-C-C-Me} \end{array}
```

IT 79637-25-5DP, conjugates with bleomycin

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of drug-delivery polymers and pharmaceutical compns. employing them)

IT 79637-25-5P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of drug-delivery polymers and pharmaceutical compns. employing
 them)

L101 ANSWER 6 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1993:537003 HCAPLUS

DN 119:137003

TI Synthesis and biological properties of polymer immunoadjuvants

AU Mao, Haiquan; Zhuo, Renxi; Fan, Changlie

CS Dep. Chem., Wuhan Univ., Wuhan, 430072, Peop. Rep. China

SO Polym. J. (Tokyo) (1993), 25(5), 499-505

CODEN: POLJB8; ISSN: 0032-3896

DT Journal

LA English

New biodegradable and biocompatible polyphosphate based on L-Try-L-Tyr or L-Ser-L-Tyr dipeptides as the recurring units were synthesized. One of these phosphates was chosen for the study of adjuvanticity in conjunction with the sol. antigen extd. from adult Schistosoma japonicum (SjAg).

Preliminary results show that, when measuring the serum antibody response to SjAg in female mice over 10 wk, one of the polyphosphates exhibited strong adjuvant activity.

IT 149513-39-3P

RL: PRP (Properties); PREP (Preparation) (prepn. and biol. properties of, as immunoadjuvant)

RN 149513-39-3 HCAPLUS

CN L-Tyrosine, N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, hexyl ester, polymer with ethyl phosphorodichloridate (9CI) (CA INDEX NAME)

CM 1

CRN 106231-84-9 CMF C32 H38 N2 O7 CDES 5:L,L

CM

CRN 1498-51-7 CMF C2 H5 C12 O2 P

ΙT 149513-39-3P 149513-45-1P 149539-41-3P

RL: PRP (Properties); PREP (Preparation) (prepn. and biol. properties of, as immunoadjuvant)

L101 ANSWER 7 OF 20 HCAPLUS COPYRIGHT 2001 ACS

1987:541105 HCAPLUS AN

DN 107:141105

ΤI Synthetic polymeric drugs

Kopecek, Jindrich; Rejmanova, Pavla; Strohalm, Jiri; Ulbrich, Karel; IN Rihova, Blanka; Chytry, Vladimir; Lloyd, John B.; Duncan, Ruth

Ceskoslovenska Akademie Ved , Czech.; Carlton Medical Products Ltd. PΑ

SO Eur. Pat. Appl., 70 pp.

CODEN: EPXXDW

DT Patent

LΑ English

FAN.	CNT	1						
	PA:	TENT NO.	KIND	DATE		APPLICATION NO.	DATE	
ΡI	ΕP	187547	A2	19860716		EP 1985-309560	19851231	<
		187547	A3	19870722				
	EΡ	187547	B1	19910227				
		R: AT, BE,	CH, DE	, FR, GB, 1	ΙΤ,	LI, LU, NL, SE		
	AT	60991	E	19910315		AT 1985-309560	19851231	<
	DK	8600033	A	19860705		DK 1986-33	19860103	<
	DK	164485	В	19920706				
	DK	164485	С	19921123				
	ΑU	8651833	A1	19860710		AU 1986-51833	19860103	<
	ΑU	589587	В2	19891019				
	CA	1305053	A1	19920714		CA 1986-498931	19860103	<
	JΡ	61243026	A2	19861029		JP 1986-137	19860104	<
	JΡ	07005474	B4	19950125				
	US	5037883	A	19910806		US 1989-438352	19891117	<
	JΡ	07300428	<b>A</b> 2	19951114		JP 1994-140759	19940531	
PRAI	GB	1985-209	19850	104 <				-
	EΡ	1985-309560		231 <				

AB A polymeric drug comprising an inert synthetic polymeric carrier combined through peptide spacers with a bioactive mol., a targeting moiety, and an

optional crosslinker, comprises (1) 5.0-99.7 mol% units derived from N-(2-hydroxypropyl)methacrylamide, (2) 0.2-20.0 mol% units derived from a N-methacryloylated peptide, the peptide groups being bound to a bioactive moiety, (3) 0.1-94.8 mol% units derived from N-methacrylamide, N-methacrylic acid, or a N-methacryloylated amino acid (peptide), to which are bound a determinant capable of interacting with specific receptors on cell surfaces, (4) optionally, 0-5 mol% units derived from a N-methacryloylated peptide, the peptide groups being bound to a linking group which is similarly attached to a similar peptide group attached to another polymer chain, and (5) optionally, as a bioassay label, 0-2 mol% units derived from N-methacryloylated tyrosinamide. Peptide groups which act as spacers in the polymer undergo lysosomal hydrolysis at a satisfactory rate to give controlled intracellular drug release. N-(2-hydroxypropyl)methacrylamide 2.5, N-methacryloylated tyrosine 0.047, N-methacryloylated Gly-Phe-Leu-Gly-O-C6H4NO2-2 0.555, and N-methacryloylated Gly-Gly-galactosamine 0.55 g were polymd. and reacted with daunomycin to give a single chain polymer contg. daunomycin bound to tetrapeptide spacer, galactosamine bound to dipeptide spacer and tyrosinamide bound directly to the main chain. Various compds. with different spacers and determinants were prepd. and tested against on L1210 mouse leukemia in vitro.

IT **79637-25-5P** 

RN 79637-25-5 HCAPLUS

CN L-Tyrosine, N-[N-[N-(2-methyl-1-oxo-2-propenyl)glycyl]-L-phenylalanyl]-, 4-nitrophenyl ester, polymer with N-(2-hydroxypropyl)-2-methyl-2-propenamide (9CI) (CA INDEX NAME)

CM 1

CRN 79637-24-4 CMF C30 H30 N4 O8 CDES 5:L,L

Absolute stereochemistry.

CM 2

CRN 21442-01-3 CMF C7 H13 N O2

```
79637-25-5P
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (prepn. of)
ΙT
     79637-25-5DP, reaction products with bleomycin
     RL: PREP (Preparation)
        (prepn. of, for intracellular drug release)
L101 ANSWER 8 OF 20 HCAPLUS COPYRIGHT 2001 ACS
AN
     1986:578286 HCAPLUS
DN
     105:178286
ΤI
     Polymers containing enzymically degradable bonds. II. Degradation of
     oligopeptide sequences connecting poly[N-(2-hydroxypropyl)methacrylamide]
     chains by lysosomal cysteine proteinases
ΑU
     Subr, V.; Kopecek, J.; Duncan, R.
CS
     Inst. Macromol. Chem., Czech. Acad. Sci., Prague, 162 06, Czech.
SO
     J. Bioact. Compat. Polym. (1986), 1(2), 133-46
     CODEN: JBCPEV; ISSN: 0883-9115
DT
     Journal
LA
     English
     Water sol. crosslinked N-(2-hydroxypropyl)methacrylamide copolymers contg.
AΒ
     15 different oligopeptide crosslinking sequences were prepd.; the relation
     between their structure and their susceptibility to hydrolysis catalyzed
     by a mixt. of lysosomal enzymes isolated from rat liver was investigated.
     The data reported provide information for the tailor-made synthesis of
     crosslinks in water sol. polymers designed as macromol. drug carriers.
IT
     79637-25-5DP, reaction products with diamines
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (prepn. and enzymic degrdn. of, drug carriers in relation to)
RN
     79637-25-5 HCAPLUS
      L-Tyrosine, \ N-[N-[N-(2-methyl-1-oxo-2-propenyl)glycyl]-L-phenylalanyl]-, \\
CN
     4-nitrophenyl ester, polymer with N-(2-hydroxypropyl)-2-methyl-2-
     propenamide (9CI) (CA INDEX NAME)
     CM
          1
     CRN
         79637-24-4
         C30 H30 N4 O8
     CMF
     CDES 5:L,L
```

# Absolute stereochemistry.

IT

CM 2

CRN 21442-01-3 C7 H13 N O2 CMF

```
O CH<sub>2</sub>
    OH
Me-CH-CH2-NH-C-C-Me
IT
     79637-25-5DP, reaction products with diamines
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (prepn. and enzymic degrdn. of, drug carriers in relation to)
ΙT
     79637-25-5P
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (prepn. of)
L101 ANSWER 9 OF 20 HCAPLUS COPYRIGHT 2001 ACS
ΑN
     1986:411957 HCAPLUS
DN
     105:11957
     The activity of complement in the presence of N-(2-
TΙ
     hydroxypropyl) methacrylamide copolymers
AU
     Simeckova, J.; Rihova, B.; Plocova, D.; Kopecek, J.
CS
     Inst. Microbiol., Czech. Acad. Sci., Prague, 142 20/4, Czech.
SO
     J. Bioact. Compat. Polym. (1986), 1(1), 20-31
     CODEN: JBCPEV; ISSN: 0883-9115
DT
     Journal
LA
     English
AΒ
     N-(2-Hydroxypropyl)methacrylamide homopolymers and copolymers contq.
     oligopeptide sequences terminated in carboxylic acid groups, amine groups,
     arom. units, or puromycin have no prominent effect on the porcine
     complement system in vitro. Inhibition of both pathways of the complement
     system occurred at concns. highly exceeding the dose suitable for
     therapeutic purposes.
IT
     79637-25-5P
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (prepn. of and complement activity response to, drug delivery systems
        in relation to)
     79637-25-5 HCAPLUS
RN
     L-Tyrosine, \ N-[N-(N-(2-methyl-1-oxo-2-propenyl)glycyl]-L-phenylalanyl]-, \\
CN
     4-nitrophenyl ester, polymer with N-(2-hydroxypropyl)-2-methyl-2-
    propenamide (9CI) (CA INDEX NAME)
    CM
    CRN
         79637-24-4
```

Absolute stereochemistry.

CDES 5:L,L

CMF C30 H30 N4 O8

CRN 21442-01-3 CMF C7 H13 N O2

$$\begin{array}{c|cccc} \text{OH} & \text{O} & \text{CH}_2 \\ | & || & || \\ \text{Me-CH-CH}_2 - \text{NH-C-C-C-Me} \end{array}$$

#### IT 79637-25-5P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of and complement activity response to, drug delivery systems
 in relation to)

L101 ANSWER 10 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1984:508627 HCAPLUS

DN 101:108627

TI Effect of the chemical structure of N-(2-hydroxypropyl)methacrylamide copolymers on their ability to induce antibody formation in inbred strains of mice

AU Rihova, B.; Kopecek, J.; Ulbrich, K.; Pospisil, M.; Mancal, P.

CS Inst. Microbiol., Czech. Acad. Sci., Prague, 142 20/4, Czech.

SO Biomaterials (Guildford, Engl.) (1984), 5(3), 143-8 CODEN: BIMADU; ISSN: 0142-9612

DT Journal

LA English

The homopolymer of N-(2-hydroxypropyl)methacrylamide (HPMA) and copolymers of HPMA differing in oligopeptide side chains [-Gly-Gly-OH; -aminocaproyl (Acap)-Phe-OH; -Acap-Leu-HMDA; -Gly-Phe-Tyr-OH] or in their content (1%, 3.5%, or 8.4% mole) of -Gly-Gly-OH side chains were investigated with respect to their ability to induce antibody formation and mitogenic reaction in inbred strains of mice. The dependence on the antigen dose, on compn. of the side chain, and on the genetic background of the immunized organism was defined. The specificity of the antibody formed was predominantly directed against oligopeptide side chains, though some part of the antibody was also produced against hydroxypropyl chains.

Neither the homopolymer nor the copolymers behaved in the tissue culture as mitogens.

IT 79637-25-5P

RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study); PREP (Preparation) (prepn. and antigenicity of, genetics of, in mouse)

RN 79637-25-5 HCAPLUS

CN L-Tyrosine, N-[N-[N-(2-methyl-1-oxo-2-propenyl)glycyl]-L-phenylalanyl]-, 4-nitrophenyl ester, polymer with N-(2-hydroxypropyl)-2-methyl-2-propenamide (9CI) (CA INDEX NAME)

CM 1

CRN 79637-24-4 CMF C30 H30 N4 O8 CDES 5:L,L

CM 2

CRN 21442-01-3 CMF C7 H13 N O2

$$\begin{array}{c|c} \text{OH} & \text{O} & \text{CH}_2 \\ & | & || & || \\ \text{Me-} & \text{CH-} & \text{CH}_2 - \text{NH-} & \text{C-} & \text{C-} & \text{Me} \end{array}$$

#### IT 79637-25-5P

RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study); PREP (Preparation) (prepn. and antigenicity of, genetics of, in mouse)

L101 ANSWER 11 OF 20 HCAPLUS COPYRIGHT 2001 ACS

1983:600416 HCAPLUS ΑN

DN 99:200416

ΤI Polymers containing enzymically degradable bonds. 8. Degradation of oligopeptide sequences in N-(2-hydroxypropyl)methacrylamide copolymers by bovine spleen cathepsin B

ΑU Rejmanova, Pavla; Kopecek, Jindrich; Pohl, Jan; Baudys, Miroslav; Kostka, Vladimir

CS Inst. Macromol. Chem., Czech. Acad. Sci., Prague, Czech.

SO Makromol. Chem. (1983), 184(10), 2009-20

CODEN: MACEAK; ISSN: 0025-116X

DT Journal

LAEnglish

AB Three types of copolymers of N-(2-hydroxypropyl)methacrylamide (HPMA) were prepd. which contain oligopeptide sequences: (a) HPMA copolymers contg. oligopeptide side-chains terminated with p-nitroaniline; (b) sol. HPMA copolymers contg. oligopeptide sequences in crosslinks connecting 2 poly(HPMA) chains; (c) a hydrophilic gel, i.e., 3-dimensional copolymer of HPMA contg. an oligopeptide sequence in the crosslinks. These polymeric substrates (suitable as drug carriers) contg. potentially degradable oligopeptide sequences were incubated with an intracellular proteolytic enzyme, bovine spleen cathepsin B [9047-22-7]. The degrdn. process of the substrates made it possible to reveal the relationship between the structure of oligopeptide sequences and their degradability. The results suggest an important role played by cathepsin B in the degrdn. of polymeric substrates investigated in this study under physiol. conditions. IT

79637-25-5DP, reaction products with amino acid or oligopeptide nitroanilides

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and degrdn. by cathepsin B, drug delivery in relation to)

RN 79637-25-5 HCAPLUS CN L-Tyrosine, N-[N-[N-(2-methyl-1-oxo-2-propenyl)glycyl]-L-phenylalanyl]-, 4-nitrophenyl ester, polymer with N-(2-hydroxypropyl)-2-methyl-2-propenamide (9CI) (CA INDEX NAME)

CM 1

CRN 79637-24-4 CMF C30 H30 N4 O8 CDES 5:L,L

Absolute stereochemistry.

CM 2

CRN 21442-01-3 CMF C7 H13 N O2

TT 79637-25-5DP, reaction products with amino acid or oligopeptide nitroanilides

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and degrdn. by cathepsin B, drug delivery in relation to)

L101 ANSWER 12 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1981:587649 HCAPLUS

DN 95:187649

TI Polymers containing enzymically degradable bonds. 2. Poly[N-(2-hydroxypropyl)methacrylamide] chains connected by oligopeptide sequences cleavable by chymotrypsin

AU Rejmanova, Pavla; Obereigner, Blahoslav; Kopecek, Jindrich

CS Inst. Macromol. Chem., Czech. Acad. Sci., Prague, 162 06, Czech.

SO Makromol. Chem. (1981), 182(7), 1899-915 CODEN: MACEAK; ISSN: 0025-116X

DT Journal

LA English

AB H2C:CMeCONHCH2CN(OH)Me was polymd. with H2C:CMeCO-X-X1-X2-OC6H4NO2-p (X = Gly, X1 = Gly, Ala, .beta.-Ala, Ile, Leu, Phe, D-Phe, Val, X2 = null; X = Ala, D-Ala, X1 = Val, X2 = null; X-X1-X2 = Gly-Gly-Phe, Gly-Gly-Val, Ala-Gly-Val, Gly-Phe-Phe, Gly-Phe-Tyr) to give the corresponding copolymers, which were crosslinked with RNH(CH2)nNHR (R = H, H-Phe, H-D-Phe, H-Tyr, H-Gly, H-Ala, H-Ala-Ala, n = 6; R = H-Gly, H-Ala, H-Ala-Gly, H-Ala-Ala, n = 2) to give title crosslinked copolymers. The latter polymers were cleaved in the oligopeptide sequences by

.alpha.-chymotrypsin. The degradability of crosslinks contg. oligopeptide sequences was dependent on both steric and structural factors, and the degradability of the crosslinks increased with increased spacing of the bond susceptible to enzymic attack from the polymer backbone.

IT 79637-25-5P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (prepn. and crosslinking of, with bis(aminoacyl)alkalinediamines)

RN 79637-25-5 HCAPLUS

CN L-Tyrosine, N-[N-[N-(2-methyl-1-oxo-2-propenyl)glycyl]-L-phenylalanyl]-, 4-nitrophenyl ester, polymer with N-(2-hydroxypropyl)-2-methyl-2-propenamide (9CI) (CA INDEX NAME)

CM 1

CRN 79637-24-4 CMF C30 H30 N4 O8 CDES 5:L,L

Absolute stereochemistry.

CM 2

CRN 21442-01-3 CMF C7 H13 N O2

IT 79637-25-5P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (prepn. and crosslinking of, with bis(aminoacyl)alkalinediamines)

IT 79637-25-5DP, reaction products with N,N'-

bis(aminoacyl)alkylenediamines

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and degrdn. of, by chymotrypsin)

L101 ANSWER 13 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1981:98801 HCAPLUS

DN 94:98801

TI Search for regular polypeptides with esterase properties. I Polypeptides containing tyrosine and glutamic acid residues

AU Shibnev, V. A.; Ismoilov, M.; Khalikov, Sh. Kh.; Lakizova, I. Yu.

CS Inst. Mol. Biol., Moscow, USSR

SO Khim. Prir. Soedin. (1979), (5), 687-99 CODEN: KPSUAR; ISSN: 0023-1150

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DT Journal
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LA Russian

The polypeptides, H-[Glu-Tyr]n-OH, H-[Glu-Tyr3]n-OH, and H-[Glu5-Tyr]-OH were enzyme-like catalysts of the hydrolysis of p-nitrophenyl acetate. The pH-activity curves for the polypeptides showed a max. in the pH range 6.9-7.2 and a much smaller max. at pH 6.0-6.2. The temp.-activity curves were bell-shaped, showing a max. at .apprx.45.degree. The Km values of the polypeptides were similar to those of chymotrypsin. CD spectra showed that at optimal pH and temp. values, the polypeptide structure was a random coil. The appearance of ordered structures (.alpha.-helix or .beta.-structure) in the polypeptide chains resulted in the decrease or complete disappearance of catalytic activity. The highest catalytic activity was obsd. with H-[Glu-Tyr]n-OH.

IT 60961-79-7

RL: BIOL (Biological study)
 (esterase activity and CD of)

RN 60961-79-7 HCAPLUS

CN L-Tyrosine, N-[N-(N-L-.alpha.-glutamyl-L-tyrosyl)-L-tyrosyl]-, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 60961-78-6 CMF C32 H36 N4 O10 CDES 5:ALL,L

Absolute stereochemistry.

#### IT 60961-79-7

RL: BIOL (Biological study)
 (esterase activity and CD of)

L101 ANSWER 14 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1978:424772 HCAPLUS

DN 89:24772

TI Synthesis of polypeptides containing tyrosine and glutamic acid residues catalyzing the hydrolysis of p-nitrophenyl acetate

AU Khalikov, Sh. Kh.; Alieva, S. V.; Ismailov, M. I.; Shibnev, V. A.

CS Tadzh. Gos. Univ., Dushanbe, USSR

SO Ref. Dokl. Soobshch. - Mendeleevsk. S'ezd Obshch. Prikl. Khim., 11th ( 1975), Volume 2, 111-12. Editor(s): Rozinskaya, V. N. Publisher: "Nauka", Moscow, USSR.

CODEN: 37MOAO

DT Conference

LA Russian

AB Polypeptides, (Tyr-Glu)n, [Tyr-(Glu)5]n, [Glu-(Tyr)3]n, [(Tyr)2-Glu-His]n, [Glu-His-Glu]n, were prepd. by polymn. of appropriate monomer peptide active esters. These polypeptides catalyzed the hydrolysis of the title acetrate.

IT 60961-79-7

CM 1

CRN 60961-78-6 CMF C32 H36 N4 O10 CDES 5:ALL,L

Absolute stereochemistry.

IT 60961-79-7

RL: RCT (Reactant)

(prepn. and nitrophenyl acetate hydrolysis in presence of)

L101 ANSWER 15 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1977:598927 HCAPLUS

DN 87:198927

TI Specificity of H-2-linked Ir gene control in mice: recognition of defined sequence analogs of (T, G)-A--L

AU Ruede, Erwin; Guenther, E.; Meyer-Delius, Margot; Liehl, E.

CS Max-Planck-Inst. Immunbiol., Freiburg/Br., Ger.

SO Eur. J. Immunol. (1977), 7(8), 520-9

CODEN: EJIMAF

DT Journal

LA English

To study the specificity of H-linked Ir gene control in more detail the AB polypeptide poly(L-Tyr, L-Glu)-poly(DL-Ala)--poly(L-Lys)((T,G)-A--L) was modified by replacing the (T,G) copolymers by defined tyrosine-contg. oligopeptides. The antibody response to all of the analogs of (T,G)-A--L tested was influenced to some extent by H-2-linked Ir gene(s), the response pattern being concordant with that of (T,G)-A--L. Some of the antigens carrying structurally and serol. distinct oligopeptides were as efficient with respect to high/low responder discrimination as (T,G)-A--L, others including the core polypeptide A--L itself were weakly immunogenic and gave only a small high-low responder split. Furthermore data indicate that in addn. to the tyrosine peptides the poly-DL-alanine side chains may play an important role for the recognition of these polypeptides. The possibility that the common response pattern could be due to an Ir gene specific mainly for the DL-alanine peptides is discussed. Specificity of genetic control would then be relatively independent of the serol. specificity of the tyrosine peptides. But, recognition of the analogs and of (T,G)-A--L could be controlled by sep. but closely linked Ir genes specific for each of the terminal peptide determinants which probably include the adjacent alanine residues.

IT 64773-07-5

RL: BIOL (Biological study)

(block, antigens, recognition of, gene H-2-linked Ir in)

RN 64773-07-5 HCAPLUS

CN L-Tyrosine, N-(N-L-.alpha.-glutamyl-L-tyrosyl)-, polymer with alanine and L-lysine (9CI) (CA INDEX NAME)

CM 1

CRN 64773-06-4 CMF C23 H27 N3 O8 CDES 5:L,L,L

Absolute stereochemistry.

CM 2

CRN 302-72-7 CMF C3 H7 N O2

CM 3

CRN 56-87-1 CMF C6 H14 N2 O2 CDES 5:L

Absolute stereochemistry.

IT 64773-07-5 64773-11-1 64773-12-2 64808-81-7 64808-82-8 64808-84-0

RL: BIOL (Biological study)

(block, antigens, recognition of, gene H-2-linked Ir in)

L101 ANSWER 16 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1977:43998 HCAPLUS

DN 86:43998

TI Synthesis of polyhexapeptide and polytetrapeptide containing residues of glutamic acid and tyrosine

AU Khalikov, Sh. Kh.; Alieva, S. V.; Ismailov, M. I.; Shibnev, V. A.

CS Tadzh. Gos. Univ. im. Lenina, Dushanbe, USSR

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SO Dokl. Akad. Nauk Tadzh. SSR (1976), 19(1), 35-9
    CODEN: DANTAL

DT Journal
LA Russian
AB [[Glu]3-Tyr]n, [Glu-[Tyr]3]n, and [Tyr-[Glu]5]n
```

AB [[Glu]3-Tyr]n, [Glu-[Tyr]3]n, and [Tyr-[Glu]5]n, possessing degrees of polymn. of 98, 43, and 35, resp., were prepd. by polymn. of the corresponding 2,4,6-Cl3C6H2 esters of o-PhCH2 blocked monomers.

IT 60961-79-7P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of)

RN 60961-79-7 HCAPLUS

CN L-Tyrosine, N-[N-(N-L-.alpha.-glutamyl-L-tyrosyl)-L-tyrosyl]-, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 60961-78-6 CMF C32 H36 N4 O10 CDES 5:ALL,L

Absolute stereochemistry.

#### IT 60961-79-7P

L101 ANSWER 17 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1976:593092 HCAPLUS

DN 85:193092

TI Some catalytic properties of regular polypeptides containing glutamic acid and tyrosine residues

AU Shibnev, V. A.; Ismoilov, M. I.

CS Inst. Mol. Biol., Moscow, USSR

SO Tezisy Dokl. - Vses. Simp. Khim. Pept. Belkov. 3rd (1974), 172 Publisher: Akad. Nauk Ukr. SSR, Kiev, USSR. CODEN: 33GEA4

DT Conference

LA Russian

AB [(Glu)m-Tyr]n (m = 1,2,3,5; n = 35-98) and [Glu-(Tyr)3]43 were prepd. by polymn. of the appropriate trichlorophenyl active ester monomers. Hydrolysis rates of AcOC6H4NO2-4 in the presence of these polymers were detd., and their catalytic activity depended on the primary structures of the polypeptides, and the pH and temp. of the hydrolysis medium.

IT 60961-79-7

RL: RCT (Reactant)

(catalyst for hydrolysis of nitrophenyl acetate)

RN 60961-79-7 HCAPLUS

CN L-Tyrosine, N-[N-(N-L-.alpha.-glutamyl-L-tyrosyl)-L-tyrosyl]-, homopolymer (9CI) (CA INDEX NAME)

CM 1

CRN 60961-78-6 CMF C32 H36 N4 O10 CDES 5:ALL,L

Absolute stereochemistry.

IT 60961-79-7

RL: RCT (Reactant)

(catalyst for hydrolysis of nitrophenyl acetate)

L101 ANSWER 18 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1976:122383 HCAPLUS

DN 84:122383

TI Grafting of amino acids having a third function onto a polymer, 1. Fixation of tyrosine onto chain ends. Model reaction

AU Gueniffey, Henri; Queromes, Armel; Garnier, Raymonde; Pinazzi, Christian

CS Lab. Chim. Org. Macromol., Le Mans, Fr.

SO Makromol. Chem. (1976), 177(2), 359-65

CODEN: MACEAK

DT Journal

LA French

The NH2 and CO2H groups of tyrosine were blocked with tosyl chloride or benzyl chloroformate and ester formation resp. and the deriv. was then reacted through its phenolic functions with .alpha.,.omega.-difunctional mols., or was dimerized and reacted with .alpha.,.omega.-difunctional mols. to form a polymer. Blocked ethyl N-(p-tosyl)tyrosinate [58559-09-4] was prepd. and reacted with hexamethylene diisocyanate (I) [822-06-0] to give diethyl 2,2'-(p-tosylamino)-3,3'-hexamethylene bis(iminocarbonyloxy-1,4-phenylene)dipropionate [58559-08-3]. The dimer ethyl N-(N-p-tosyltyrosyl)tyrosinate [58557-95-2] was prepd. from ethyl tyrosinate chlorohydrate [4089-07-0] and condensed with I to form a polymer.

IT 58557-94-1P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of)

RN 58557-94-1 HCAPLUS

CN L-Tyrosine, N-[N-[(phenylmethoxy)carbonyl]-L-tyrosyl]-, ethyl ester, polymer with 1,6-diisocyanatohexane (9CI) (CA INDEX NAME)

CM 1

CRN 4142-95-4 CMF C28 H30 N2 O7 CDES 5:L,L

CM 2

CRN 822-06-0 CMF C8 H12 N2 O2

OCN-(CH<sub>2</sub>)<sub>6</sub>-NCO

## IT 58557-94-1P 58557-96-3P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of)

L101 ANSWER 19 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1976:106037 HCAPLUS

DN 84:106037

TI Synthesis of regular polypeptides containing residual tyrosine and glutamic acid

AU Shibnev, V. A.; Ismoilov, M. I.; Khalikov, Sh. Kh.

CS Inst. Mol. Biol., Moscow, USSR

SO Izv. Akad. Nauk SSSR, Ser. Khim. (1975), (9), 2082-8 CODEN: IASKA6

DT Journal

LA Russian

IT 58436-99-0P

RN 58436-99-0 HCAPLUS

CN L-Tyrosine, N-[N-L-.alpha.-glutamyl-O-(phenylmethyl)-L-tyrosyl]-O-(phenylmethyl)-, 5-(phenylmethyl) 1-(2,4,5-trichlorophenyl) ester, monohydrochloride, homopolymer (9CI) (CA INDEX NAME)

CM 1 ·

CRN 58436-98-9 CMF C50 H46 C13 N3 O8 . C1 H CDES 5:L,L,L

PAGE 2-A

HCl

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IT 58436-99-0P 58437-05-1P
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L101 ANSWER 20 OF 20 HCAPLUS COPYRIGHT 2001 ACS

AN 1973:442897 HCAPLUS

DN 79:42897

TI Optically active polymers. XI. Phenol-formaldehyde polycondensates of N-benzoyl-L-tyrosine (TyB) and of N-(p-hydroxyphenylsulfonyl)-L-phenylalanine (PhHPS). Synthesis and optical activity.

AU Beaumais, Jacques; Vert, Michel; Selegny, Eric

CS U.E.R. Sci. Exactes Nat., Univ. Rouen, Mont-Saint-Aignan, Fr.

SO Makromol. Chem. (1973), 165, 17-29

CODEN: MACEAK

DT Journal

LA French

AB Acidic and basic polycondensation of formaldehyde [50-00-0] with the optically active N-benzoyl-L-tyrosine (I) and N-p-hydroxyphenylsulfonyl-L-phenylalanine gave 4 polymers characterized by ir spectra, mol. wt., comparison with similar polycondensates, and ORD of the K salts. The optical rotatory activities of acidic- and basic-prepn. media of N-benzoyl-L-tyrosine-formaldehyde copolymer [41034-34-8] differed and varied with solvent. The polymers from I and alk.-prepd. formaldehyde-N-(p-hydroxyphenylsulfonyl)-L-phenylalanine copolymer [41034-35-9] had the expected chem. structures, however, the acidic phenylalanine underwent a secondary reaction to give poly(N-(p-hydroxyphenylsulfonyl)-3-carboxy-1,2,3,4-tetrahydroisoquinoline] (I) [41026-00-0].

IT 41034-34-8P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and optical activity of)

RN 41034-34-8 HCAPLUS

CN L-Tyrosine, N-benzoyl-, polymer with formaldehyde (9CI) (CA INDEX NAME)

CM 1

CRN 2566-23-6

CMF C16 H15 N O4 CDES 5:L

Absolute stereochemistry.

CM 2

CRN 50-00-0 CMF C H2 O

 $H_2C = O$ 

#### IT 41034-34-8P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. and optical activity of)

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		219622-84-1	REGISTRY
96	RN	191858-74-9	REGISTRY
DR	223114-13-4		
97	RN	191858-72-7	REGISTRY
DR	223114-11-2		
98	RN	191858-70-5	REGISTRY
DR	223114-15-6		
99	RN	191858-68-1	REGISTRY
DR	223114-10-1		
100	RN	189760-16-5	REGISTRY
101	RN	189760-14-3	REGISTRY
102	RN	189760-12-1	REGISTRY
103	RN	189760-11-0	REGISTRY
104	RN	189760-09-6	REGISTRY
105	RN	189760-09-6	
106	RN		REGISTRY
		188716-30-5	REGISTRY
107	RN	188713-11-3	REGISTRY
108	RN	188713-10-2	REGISTRY
109	RN	188713-09-9	REGISTRY
110	RN	188713-08-8	REGISTRY
111	RN	188713-05-5	REGISTRY
112	RN	188713-03-3	REGISTRY
113	RN	188713-01-1	REGISTRY
114	RN	188712-99-4	REGISTRY
115	RN	188712-97-2	REGISTRY
116	RN	188712-95-0	REGISTRY
117	RN	188712-92-7	REGISTRY
118	RN	183480-55-9	REGISTRY
DR	188709-60-6,	188711-38-8	
119	RN	183480-54-8	REGISTRY
DR	188709-59-3		1.20101111
120	RN	183480-53-7	REGISTRY
DR	188709-58-2,	188711-36-6	REGIETA
121	RN	183480-51-5	REGISTRY
122	RN	183480-50-4	REGISTRY
123	RN	183480-48-0	
124	RN	174702-87-5	REGISTRY
125	RN		REGISTRY
126		174702-86-4	REGISTRY
	RN	174702-85-3	REGISTRY
127	RN	171436-78-5	REGISTRY
128	RN	171436-77-4	REGISTRY
129	RN	171436-76-3	REGISTRY
130	RN	171436-75-2	REGISTRY
131	RN	149826-02-8	REGISTRY
132	RN	149787-42-8	REGISTRY
133	RN	149787-41-7	REGISTRY
134	RN .	149787-40-6	REGISTRY
135	RN	149787-39-3	REGISTRY
136	RN	149787-38-2	REGISTRY
137	RN	143715-03-1	REGISTRY
138	RN	133418-81-2	REGISTRY
139	RN	133134-42-6	REGISTRY
	= := *	200101 42 0	

<sup>=&</sup>gt; d 180 ide can 1-3 10 20 30 40 50 60 70 80 90 93 96 100 106 108 114 118 124 127 131 132 137-139

L80 ANSWER 1 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 319916-60-4 REGISTRY

Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-[(1-methylethoxy)carbonyl]-1,2-CN ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

MF (C22 H23 N O6) n

CI PMS

PCT Polyamide, Polycarbonate

SR CA

LC STN Files: CA, CAPLUS, TOXLIT

1 REFERENCES IN FILE CA (1967 TO DATE) 1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 134:105786

ANSWER 2 OF 139 REGISTRY COPYRIGHT 2001 ACS L80

RN263565-88-4 REGISTRY

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, ethyl ester, polymer CNwith .alpha.-carboxy-.omega.-(carboxyoxy)poly(oxy-1,2-ethanediyl) (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

Poly(oxy-1,2-ethanediyl), .alpha.-carboxy-.omega.-(carboxyoxy)-, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine ethyl ester (9CI)

FS STEREOSEARCH

MF (C20 H23 N O5 . (C2 H4 O)n C2 H2 O5) $\mathbf{x}$ 

CI-PMS-

PCT Polyamide, Polyamide formed, Polycarbonate, Polycarbonate formed, Polyester, Polyester formed, Polyether

SR

LÇ STN Files: CA, CAPLUS, TOXLIT

> CM1

CRN 135313-59-6 CMF C20 H23 N O5

Absolute stereochemistry.

CM 2

CRN 85022-96-4

(C2 H4 O)n C2 H2 O5 CMF

CCI **PMS** 

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 132:270023

L80 ANSWER 3 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247077-80-1 REGISTRY

CN L-Tyrosine, N-[(4-hydroxyphenyl)acetyl]-, octyl ester, polymer with decanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Decanedioic acid, polymer with N-[(4-hydroxyphenyl)acetyl]-L-tyrosine octyl ester (9CI)

FS STEREOSEARCH

MF (C25 H33 N O5 . C10 H18 O4) x

CI PMS

PCT Polyamide, Polyester, Polyester formed

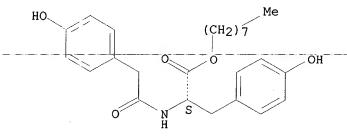
SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 240418-76-2 CMF C25 H33 N O5

Absolute stereochemistry.



CM 2

CRN 111-20-6 CMF C10 H18 O4

 $HO_2C^-$  (CH<sub>2</sub>)<sub>8</sub> - CO<sub>2</sub>H

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

L80 ANSWER 10 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247077-73-2 REGISTRY

CN L-Tyrosine, N-[(4-hydroxyphenyl)acetyl]-, octyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Butanedioic acid, polymer with N-[(4-hydroxyphenyl)acetyl]-L-tyrosine octyl ester (9CI)

FS STEREOSEARCH

MF (C25 H33 N O5 . C4 H6 O4)x

CI PMS

PCT Polyamide, Polyester, Polyester formed

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 240418-76-2 CMF C25 H33 N O5

Absolute stereochemistry.

CM 2

CRN 110-15-6 CMF C4 H6 O4

 $HO_2C-CH_2-CH_2-CO_2H$ 

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES\_IN\_FILE\_CAPLUS\_(1967\_TO\_DATE)\_\_\_\_\_

REFERENCE 1: 131:299854

L80 ANSWER 20 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247077-63-0 REGISTRY

CN L-Tyrosine, N-[(4-hydroxyphenyl)acetyl]-, ethyl ester, polymer with 2,2'-[1,2-ethanediylbis(oxy)]bis[acetic acid] (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES:

CN Acetic acid, 2,2'-[1,2-ethanediylbis(oxy)]bis-, polymer with N-[(4-hydroxyphenyl)acetyl]-L-tyrosine ethyl ester (9CI)

FS STEREOSEARCH

MF (C19 H21 N O5 . C6 H10 O6)x

CI PMS

PCT Polyamide, Polyester, Polyester formed, Polyether

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 240418-74-0 CMF C19 H21 N O5

CM 2

CRN 23243-68-7 CMF C6 H10 O6

 ${\tt HO_2C-CH_2-O-CH_2-CH_2-O-CH_2-CO_2H}$ 

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

L80 ANSWER 30 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247077-50-5 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, phenylmethyl ester, polymer with hexanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Hexanedioic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine phenylmethyl ester (9CI)

FS STEREOSEARCH

MF (C25 H25 N O5 . C6 H10 O4) $\times$ 

CI\_\_PMS

PCT Polyamide, Polyester, Polyester formed

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 189760-08-5 CMF C25 H25 N O5

Absolute stereochemistry.

CM 2

CRN 124-04-9 CMF C6 H10 O4  $HO_2C-(CH_2)_4-CO_2H$ 

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

L80 ANSWER 40 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247077-33-4 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, 1-methylpropyl ester, polymer with pentanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Pentanedioic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine 1-methylpropyl ester (9CI)

FS STEREOSEARCH

MF (C22 H27 N O5 . C5 H8 O4)x

CI PMS

PCT Polyamide, Polyester, Polyester formed

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 240418-73-9 CMF C22 H27 N O5

Absolute stereochemistry.

CM 2

CRN 110-94-1 CMF C5 H8 O4

 $HO_2C-(CH_2)_3-CO_2H$ 

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

L80 ANSWER 50 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247077-12-9 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, 1-methylethyl ester, polymer with 2,2'-[1,2-ethanediylbis(oxy)]bis[acetic acid] (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Acetic acid, 2,2'-[1,2-ethanediylbis(oxy)]bis-, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine 1-methylethyl ester (9CI) FS STEREOSEARCH

MF (C21 H25 N O5 . C6 H10 O6) x

CI PMS

PCT Polyamide, Polyester, Polyester formed, Polyether

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 189760-10-9 CMF C21 H25 N O5

# Absolute stereochemistry.

CM 2

CRN 23243-68-7 CMF C6 H10 O6

 $HO_2C-CH_2-O-CH_2-CH_2-O-CH_2-CO_2H$ 

1 REFERENCES IN FILE CA (1967 TO DATE)

1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

L80 ANSWER 60 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247076-92-2 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, 2-(2-ethoxyethoxy)ethyl ester, polymer with hexanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Hexanedioic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine 2-(2-ethoxyethoxy)ethyl ester (9CI)

FS STEREOSEARCH

MF (C24 H31 N O7 . C6 H10 O4) $\times$ 

CI PMS

PCT Polyamide, Polyester, Polyester formed

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 189760-06-3 CMF C24 H31 N O7

CRN 124-04-9 CMF C6 H10 O4

 $HO_2C-(CH_2)_4-CO_2H$ 

1 REFERENCES IN FILE CA (1967 TO DATE)

1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

L80 ANSWER 70 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247076-81-9 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, octyl ester, polymer with 3-methylhexanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Hexanedioic acid, 3-methyl-, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine octyl ester (9CI)

FS STEREOSEARCH

MF (C26 H35 N O5 . C7 H12 O4) $\times$ 

CI PMS

PCT Polyamide, Polyamide formed, Polyester, Polyester formed

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 135313-58-5 CMF C26 H35 N O5

Absolute stereochemistry.

CM 2

CRN 3058-01-3 CMF C7 H12 O4  $\begin{array}{c} & \text{Me} \\ & | \\ \text{HO}_2\text{C} - \text{CH}_2 - \text{CH} - \text{CH}_2 - \text{CH}_2 - \text{CO}_2\text{H} \end{array}$ 

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

L80 ANSWER 80 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247076-71-7 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, butyl ester, polymer with pentanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Pentanedioic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine butyl ester (9CI)

FS STEREOSEARCH

MF (C22 H27 N O5 . C5 H8 O4) $\times$ 

CI PMS

PCT Polyamide, Polyamide formed, Polyester, Polyester formed

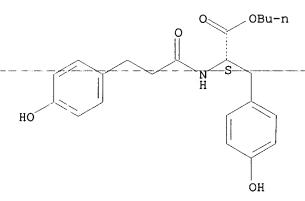
SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 174702-84-2 CMF C22 H27 N O5

Absolute stereochemistry.



CM 2

CRN 110-94-1 CMF C5 H8 O4

 $HO_2C-(CH_2)_3-CO_2H$ 

1 REFERENCES IN FILE CA (1967 TO DATE)

1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

L80 ANSWER 90 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 247076-61-5 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, methyl ester, polymer with 2,2'-oxybis[acetic acid] (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

Acetic acid, 2,2'-oxybis-, polymer with N-[3-(4-hydroxyphenyl)-1-CN oxopropyl]-L-tyrosine methyl ester (9CI)

FS STEREOSEARCH

(C19 H21 N O5 . C4 H6 O5)xMF

CI **PMS** 

PCT Polyamide, Polyester, Polyester formed, Polyether

SR

LC STN Files: CA, CAPLUS

> 1 CM

CRN 240418-72-8 CMF C19 H21 N O5

Absolute stereochemistry.

CM

CRN 110-99-6 CMF C4 H6 O5

HO2C-СH2-О-СH2-СО2Н

1 REFERENCES IN FILE CA (1967 TO DATE) 1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

L80 ANSWER 93 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 219622-86-3 REGISTRY

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, ethyl ester, polymer CN with decanedicyl dichloride (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Decanedicyl dichloride, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-Ltyrosine ethyl ester (9CI)

FS STEREOSEARCH

MF (C20 H23 N O5 . C10 H16 C12 O2)x

CI

PCT Polyamide, Polyamide formed, Polyester, Polyester formed

SR

LC STN Files: CA, CAPLUS

> CM 1

135313-59-6 CRN C20 H23 N O5 CMF

CRN 111-19-3 CMF C10 H16 C12 O2

1 REFERENCES IN FILE CA (1967 TO DATE)

1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 130:110770

L80 ANSWER 96 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 191858-74-9 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic dichloride and .alpha.-hydro-.omega.-hydroxypoly(oxy-1,2-ethanediyl), block (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Carbonic dichloride, polymer with .alpha.-hydro-.omega.-hydroxypoly(oxy-1,2-ethanediyl) and N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine hexyl ester, block (9CI)

CN Poly(oxy-1,2-ethanediyl), .alpha.-hydro-.omega.-hydroxy-, polymer with carbonic dichloride and N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine hexyl ester, block (9CI)

FS STEREOSEARCH

DR 223114-13-4

MF (C24 H31 N O5 . (C2 H4 O) n H2 O . C C12 O) x

CI PMS

PCT Polyamide, Polyamide formed, Polycarbonate, Polycarbonate formed, Polyester, Polyester formed, Polyether

SR CA

LC STN Files: CA, CAPLUS, USPATFULL

CM 1

CRN 133063-33-9 CMF C24.H31 N O5

CRN 25322-68-3

CMF (C2 H4 O)n H2 O

CCI PMS

$$\begin{array}{c|c} \text{HO} & \hline & \text{CH}_2\text{--}\text{CH}_2\text{--}\text{O} & \hline & \text{H} \end{array}$$

CM 3

CRN 75-44-5 CMF C C12 O

1 REFERENCES IN FILE CA (1967 TO DATE)

1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 127:82246

L80 ANSWER 100 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 189760-16-5 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, dodecyl ester, polymer with 2,2'-oxybis[acetic acid] (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Acetic acid, 2,2'-oxybis-, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine dodecyl ester (9CI)

FS STEREOSEARCH

MF (C30 H43 N O5 . C4 H6 O5)x

CI PMS

PCT Polyamide, Polyester, Polyester formed, Polyether

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 189760-15-4 CMF C30 H43 N O5

CRN 110-99-6 CMF C4 H6 O5

HO2C-CH2-O-CH2-CO2H

2 REFERENCES IN FILE CA (1967 TO DATE)
2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 131:299854

REFERENCE 2: 126:330869

L80 ANSWER 106 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 188716-30-5 REGISTRY

CN Poly[oxycarbonyloxy-1,4-phenylene[2-[(phenylmethoxy)carbonyl]-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene], (S)- (9CI) (CA INDEX NAME)

MF (C26 H23 N O6) n

CI PMS

PCT Polyamide, Polycarbonate

SR CA

LC STN Files: CA, CAPLUS, TOXLIT

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 126:255430

L80 ANSWER 108 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 188713-10-2 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, octyl ester, polymer with octanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Octanedioic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine octyl ester (9CI)

FS STEREOSEARCH

MF (C26 H35 N O5 . C8 H14 O4)x

CI PMS

PCT Polyamide, Polyamide formed, Polyester, Polyester formed

SR CF

LC STN Files: CA, CAPLUS, TOXLIT

CM 1

CRN 135313-58-5 CMF C26 H35 N O5

Absolute stereochemistry.

Me 
$$(CH_2)_{7}$$
 O O O OH

CM 2

CRN 505-48-6 CMF C8 H14 O4

 $HO_2C-(CH_2)_6-CO_2H$ 

3 REFERENCES IN FILE CA (1967 TO DATE)

3 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 132:348250

REFERENCE 2: 131:299854

REFERENCE 3: 126:255427

L80 ANSWER 114 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 188712-99-4 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, butyl ester, polymer with butanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Butanedioic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine butyl ester (9CI)

FS STEREOSEARCH

MF (C22 H27 N O5 . C4 H6 O4)  $\times$ 

CI PMS

PCT Polyamide, Polyamide formed, Polyester, Polyester formed

SR CA

LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL

CM 1

CRN 174702-84-2 CMF C22 H27 N O5

CRN 110-15-6 CMF C4 H6 O4

 ${\tt HO_2C-CH_2-CH_2-CO_2H}$ 

5 REFERENCES IN FILE CA (1967 TO DATE) 5 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 133:168440

REFERENCE 2: 132:348250

REFERENCE 3: 131:299854

REFERENCE 4: 129:306444

REFERENCE 5: 126:255427

L80 ANSWER 118 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 183480-55-9 REGISTRY

CN Poly[oxycarbonyloxy-1,4-phenylene[(2S)-2-[(octyloxy)carbonyl]-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Poly[oxycarbonyloxy-1,4-phenylene[2-[(octyloxy)carbonyl]-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene], (S)-

DR 188709-60-6, 188711-38-8

MF (C27 H33 N O6)n

CI PMS

PCT Polyamide, Polycarbonate

SR CF

LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL

6 REFERENCES IN FILE CA (1967 TO DATE) 6 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 134:105785

REFERENCE 2: 132:212631

REFERENCE 3: 126:255433

REFERENCE 4: 126:255430

REFERENCE 5: 126:255428

REFERENCE 6: 126:8702

L80 ANSWER 124 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 174702-87-5 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, octyl ester, polymer with carbonic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Carbonic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine octyl ester (9CI)

FS STEREOSEARCH

MF (C26 H35 N O5 . C H2 O3) $\times$ 

CI PMS

PCT Polyamide, Polyamide formed, Polycarbonate, Polycarbonate formed,

Polyester, Polyester formed

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 135313-58-5 CMF C26 H35 N O5

Absolute stereochemistry.

CM 2

CRN 463-79-6 CMF C H2 O3

О || НО— С— ОН

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 124:203129

L80 ANSWER 127 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 171436-78-5 REGISTRY

CN Poly[oxycarbonyloxy-1,4-phenylene[2-[(octyloxy)carbonyl]-1,2-ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene] (9CI) (CA INDEX NAME)

MF (C27 H33 N O6)n

CI PMS

PCT Polyamide, Polycarbonate

SR CA

LC STN Files: CA, CAPLUS, TOXLIT

2 REFERENCES IN FILE CA (1967 TO DATE)
2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 124:211936

REFERENCE 2: 124:15445

L80 ANSWER 131 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 149826-02-8 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, octyl ester, polymer with hexanedioic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Hexanedioic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-Ltyrosine\_octyl\_ester\_(9CI)\_\_\_\_\_

FS STEREOSEARCH

MF (C26 H35 N O5 . C6 H10 O4)  $\times$ 

CI PMS

PCT Polyamide, Polyamide formed, Polyester, Polyester formed

SR CA

LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL

CM 1

CRN 135313-58-5 CMF C26 H35 N O5

CRN 124-04-9 CMF C6 H10 O4

## $HO_2C-(CH_2)_4-CO_2H$

8 REFERENCES IN FILE CA (1967 TO DATE)

8 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 132:348250

REFERENCE 2: 132:313446

REFERENCE 3: 131:299854

REFERENCE 4: 129:306444

REFERENCE 5: 126:255427

REFERENCE 6: 124:37506

REFERENCE 7: 121:163954

REFERENCE 8: 119:146603

L80 ANSWER 132 OF 139 REGISTRY COPYRIGHT 2001 ACS

RN 149787-42-8 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with 1,4-benzenedicarboxylic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 1,4-Benzenedicarboxylic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine hexyl ester (9CI)

FS STEREOSEARCH

MF (C24 H31 N O5 . C8 H6 O4)x

CI PMS

PCT \_Polyamide, Polyamide\_formed, Polyester, Polyester formed\_\_\_\_

SR CA

LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL

CM 1

CRN 133063-33-9 CMF C24 H31 N O5

## Absolute stereochemistry.

CM 2

CRN 100-21-0 CMF C8 H6 O4

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 119:146603

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RN 143715-03-1 REGISTRY

CN L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer with carbonic acid (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Carbonic acid, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-tyrosine hexyl ester (9CI)

FS STEREOSEARCH

MF (C24 H31 N O5 . C H2 O3) $\times$ 

CI PMS

PCT Polyamide, Polyamide formed, Polycarbonate, Polycarbonate formed, Polyester, Polyester formed

SR CA

LC STN Files: CA, CAPLUS, TOXLIT

CM 1

CRN 133063-33-9 CMF C24 H31 N O5

Absolute stereochemistry.

CM 2

CRN 463-79-6 CMF C H2 O3

2 REFERENCES IN FILE CA (1967 TO DATE)
2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 124:203129

REFERENCE 2: 117:157568

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133418-81-2 REGISTRY RN

L-Tyrosine, N-[3-(4-hydroxyphenyl)-1-oxopropyl]-, hexyl ester, polymer CN with carbonic dichloride (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

Carbonic dichloride, polymer with N-[3-(4-hydroxyphenyl)-1-oxopropyl]-L-CN tyrosine hexyl ester (9CI)

STEREOSEARCH FS

(C24 H31 N O5 . C C12 O)x MF

**PMS** CI

Polyamide, Polyamide formed, Polycarbonate, Polycarbonate formed, PCT Polyester, Polyester formed

SR

CA, CAPLUS, CHEMCATS, TOXLIT, USPATFULL STN Files: LC

CM

133063-33-9 CRN C24 H31 N O5 CMF

Absolute stereochemistry.

2 CM

CRN 75-44-5 C C12 O CMF

4 REFERENCES IN FILE CA (1967 TO DATE)

4 REFERENCES IN FILE CAPLUS (1967 TO DATE)

126:8702 1: REFERENCE

2: 117:70492 REFERENCE

116:201112 REFERENCE 3:

REFERENCE 114:186204 4:

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133134-42-6 REGISTRY RN

Poly[oxycarbonimidoyloxy-1,4-phenylene[2-[(hexyloxy)carbonyl]-1,2-CN ethanediyl]imino(1-oxo-1,3-propanediyl)-1,4-phenylene], (S)- (9CI) (CA INDEX NAME)

MF (C25 H30 N2 O5)n

CI **PMS** 

PCT Polyamide

SR CA

CA, CAPLUS, TOXLIT LC STN Files:

1 REFERENCES IN FILE CA (1967 TO DATE) 1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 114:171127